2x1 con

Graser 10/030313 Applicant

=> d his

(FILE 'HOME' ENTERED AT 11:24:56 ON 16 DEC 2004)

FILE 'HCAPLUS' ENTERED AT 11:25:27 ON 16 DEC 2004

E W02000-CA811/AP, PRN

L1 1 WO2000-CA811/AP, PRN

L2 1 US6391313/PN

1 L1-2

FILE 'REGISTRY' ENTERED AT 11:26:35 ON 16 DEC 2004

FILE 'HCAPLUS' ENTERED AT 11:26:37 ON 16 DEC 2004
TRA L3 1- RN : 4 TERMS

FILE 'REGISTRY' ENTERED AT 11:26:37 ON 16 DEC 2004

L5 4 SEA L4

FILE 'WPIX' ENTERED AT 11:26:42 ON 16 DEC 2004

E WO2000-CA811/AP,PRN

L6 1 WO2000-CA811/AP, PRN

L7 1 US6391313/PN

L8 1 L6-7

=> b hcap

FILE 'HCAPLUS' ENTERED AT 11:27:25 ON 16 DEC 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 16 Dec 2004 VOL 141 ISS 25 FILE LAST UPDATED: 15 Dec 2004 (20041215/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

## => d all 13

L3 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2001:63846 HCAPLUS

DN 134:120915

ED Entered STN: 26 Jan 2001

TI Multicomponent vaccine to protect against disease caused by Haemophilus influenzae and Moraxella catarrhalis

IN Loosmore, Sheena M.; Yang, Yan-Ping; Klein, Michel H.; Sasaki, Ken

PA Connaught Laboratories Limited, Can.

SO PCT Int. Appl., 58 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K039-00

CC 63-3 (Pharmaceuticals)

Section cross-reference(s): 15

FAN. CNT 1

1.774.	-111	-																		
	PATENT NO.						D	DATE			APPLICATION NO.					D				
							-					- <b></b>				-				
ΡI	WO 2001005424					A2	A2 20010125			1	WO 2000-CA811						20000711 <			
	WO	2001005424			<b>A3</b>		20010802													
		W:	ΑE,	AG,				AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,		
			CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,		
			HU,	ID,	IL,	IN,	IS,	ĴΡ,	KE,	KG,	KP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,		
			LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PL,	PT,	RO,	RU,		
			SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	TZ,	UA,	ŪĠ,	US,	UΖ,	VN,		
			YU,	ZA,	ZW,	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM						

Search done by Noble Jarrell

Joseph Johnson

ophias 1

. .

```
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     US 6391313
                             В1
                                    20020521
                                                 US 1999-353617
                                                                            19990715 <--
                                    20010125
                                                 CA 2000-2378862
                                                                            20000711 <--
     CA 2378862
                             AA
     EP 1200122
                             A2
                                    20020502
                                                 EP 2000-945494
                                                                           20000711 <--
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, SI, LT, LV, FI, RO, MK, CY, AL
     AU 767096
                            В2
                                    20031030
                                                 AU 2000-59586
                                                                            20000711 <--
PRAI US 1999-353617
                                    19990715
                             Α
     WO 2000-CA811
                             W
                                    20000711
CLASS
PATENT NO.
                   CLASS PATENT FAMILY CLASSIFICATION CODES
                          A61K039-00
WO 2001005424
                   TCM
US 6391313
                   ECLA
                          A61K039/116
AB A multi-valent immunogenic composition confers protection on an immunized host
     against infection caused by both Haemophilus influenzae and Moraxella
     catarrhalis. Such composition comprises at least four antigens comprising at
     least one antigen from Haemophilus influenzae, and at least one antigen
     from Moraxella catarrhalis. Three of the antigens are adhesins. High
     mol. weight (HMW) proteins and Haemophilus influenzae adhesin (Hia) proteins
     of non-typeable Haemophilus and a 200 kDa outer membrane protein of
     Moraxella catarrhalis comprise the adhesin components while the other
     antigen is a non-proteolytic analog of Hin47 protein. Each component does not impair the immunogenicity of the others. The multi-valent immunogenic composition may be combined with DTP component vaccines, which may also include
     non-virulent poliovirus and PRP-T, to provide a component vaccine without
     impairment of the immunogenic properties of the other antigens.
     adhesin antigen vaccine Haemophilus Moraxella
     Proteins, specific or class
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
         (HMW1; multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
     Proteins, specific or class
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
         (HMW2; multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
     Proteins, specific or class
IT
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
         (Hin47; multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
     Proteins, specific or class
IT
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
         (Hsf (Haemophilus surface fibril); multicomponent vaccine to protect
         against disease caused by Haemophilus influenzae and Moraxella
         catarrhalis)
IT
     Proteins, specific or class
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
         (OMP (outer membrane protein); multicomponent vaccine to protect
         against disease caused by Haemophilus influenzae and Moraxella
         catarrhalis)
IT
     Immunostimulants
         (adjuvants; multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
IT
     Agglutinins and Lectins
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (agglutinogens; multicomponent vaccine to protect against disease
```

ROUT

caused by Haemophilus influenzae and Moraxella catarrhalis)

Adhesins

TT

```
RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); PEP (Physical, engineering or Chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
         (antigenic; multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
TT
     Toxoids
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (diphtheria; multicomponent vaccine to protect against disease caused
         by Haemophilus influenzae and Moraxella catarrhalis)
     Organelle
         (fibril, surface; multicomponent vaccine to protect against disease
         caused by Haemophilus influenzae and Moraxella catarrhalis)
IT
     Hemagglutinins
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU
     (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (filamentous; multicomponent vaccine to protect against disease caused
         by Haemophilus influenzae and Moraxella catarrhalis)
     Chinchilla
IT
     Haemophilus influenzae
     Molecular cloning
     Molecular weight distribution
     Moraxella catarrhalis
     Polyacrylamide gel electrophoresis
     Vaccines
         (multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
IT
     Antigens
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
         (multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
     Heat-shock proteins
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
         (non-proteolytic; multicomponent vaccine to protect against disease
         caused by Haemophilus influenzae and Moraxella catarrhalis)
IT
     Human poliovirus
         (non-virulent; multicomponent vaccine to protect against disease caused
         by Haemophilus influenzae and Moraxella catarrhalis)
IT
     Ear
         (otitis, otitis media; multicomponent vaccine to protect against
         disease caused by Haemophilus influenzae and Moraxella catarrhalis)
IT
     Agglutinins and Lectins
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (pertactins; multicomponent vaccine to protect against disease caused
         by Haemophilus influenzae and Moraxella catarrhalis)
IT
     Toxoids
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (pertussis; multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
ΙT
     Mutation
         (substitution; multicomponent vaccine to protect against disease caused
         by Haemophilus influenzae and Moraxella catarrhalis)
IT
     Toxoids
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (tetanus; multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
     9001-92-7, Proteinase
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); BIOL (Biological
     study); OCCU (Occurrence)
```

(activity levels; multicomponent vaccine to protect against disease caused by Haemophilus influenzae and Moraxella catarrhalis) 7784-30-7, Aluminum phosphate 21645-51-2, Aluminum hydroxide, biological IT studies RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (adjuvant; multicomponent vaccine to protect against disease caused by Haemophilus influenzae and Moraxella catarrhalis) 151-21-3, Sds, analysis RL: ARU (Analytical role, unclassified); ANST (Analytical study) (multicomponent vaccine to protect against disease caused by Haemophilus influenzae and Moraxella catarrhalis) FILE 'REGISTRY' ENTERED AT 11:27:31 ON 16 DEC 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 American Chemical Society (ACS) Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem. STRUCTURE FILE UPDATES: 14 DEC 2004 HIGHEST RN 797749-23-6 DICTIONARY FILE UPDATES: 14 DEC 2004 HIGHEST RN 797749-23-6 TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004 Please note that search-term pricing does apply when conducting SmartSELECT searches. Crossover limits have been increased. See HELP CROSSOVER for details. Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html => d ide 15 tot ANSWER 1 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN 21645-51-2 REGISTRY Aluminum hydroxide (Al(OH)3) (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES: Aluminum hydroxide (6CI, 8CI) OTHER NAMES: 42STE CN CN A 3011 AC 400 CN CNAC 400 (hydroxide) CNAC 450 CN AC 714KC AE 107 CN AF 260 CNCN AKP-DA CNAlcan SF 4 Alcoa 331 CN CN Alcoa 710 CN Alcoa A 325 CN Alcoa AS 301 Alcoa C 30BF Alcoa C 31 CN CN CNAlcoa C 33 Alcoa C 330 CN CN Alcoa C 331 Alcoa C 333 CN Alcoa C 385 CN Alcoa H 65 CN CN Alcoa OC 1000 CN Alhydrogel

CN

CN

CN CN

CN

Alolt 50AF

Alolt 59 Alolt 60FLS

Alolt 8 Alolt 80

```
CN
     Alolt 90
     Alternagel
CN
     Alugel
CN
     Alugelibys
CN
CN
     Alumigel
CN
     Alumina trihydrate
     Aluminic acid (H3AlO3)
CN
CN
      Aluminum oxide (Al2O3), trihydrate
CN
     Aluminum oxide trihydrate
     Aluminum trihydroxide
CN
CN
     Alusal
CN
     Amberol ST 140F
CN
     Amphogel
     Amphoiel
CN
     Antipollon HT
CN
CN
     Apyral
CN
     Apyral 120
     Apyral 120VAW
CN
CN
     Apyral 15
CN
     Apyral 2
ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for
     546141-62-2, 12252-70-9, 13783-16-9, 8012-63-3, 8064-00-4, 1302-29-0, 128083-27-2, 106152-09-4, 51330-22-4, 151393-94-1, 159704-77-5
DR
MF
     Al H3 O3
ÇI
     COM
                   ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
LC
     STN Files:
        CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
        CHEMLIST, CHEMSAFE, CIN, CSCHEM, CSNB, DDFU, DETHERM*, DIOGENES, DIPPR*,
        DRUGU, EMBASE, ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT, ENCOMPPAT2, GMELIN*,
        HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC, PDLCOM*, PIRA, PROMT, PS, RTECS*, TOXCENTER, TULSA, USAN, USPAT2,
        USPATFULL, VETU, VTB
          (*File contains numerically searchable property data)
      Other Sources: DSL**, EINECS**, TSCA**
          (**Enter CHEMLIST File for up-to-date regulatory information)
DT.CA CAplus document type: Book; Conference; Dissertation; Journal; Patent;
        Roles from patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU
RI. P
        (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
(Reactant or reagent); USES (Uses); NORL (No role in record)
RLD.P Roles for non-specific derivatives from patents: ANST (Analytical
        study); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation);
        PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES
RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); CMBI (Combinatorial study); FORM (Formation, nonpreparative);
        MSC (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC
        (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses);
        NORL (No role in record)
RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical
        study); BIOL (Biological study); FORM (Formation, nonpreparative); MSC
        (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process);
        PRP (Properties); RACT (Reactant or reagent); USES (Uses)
     OH
HO-A1-OH
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
             22458 REFERENCES IN FILE CA (1907 TO DATE)
               374 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            22498 REFERENCES IN FILE CAPLUS (1907 TO DATE)
                 1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)
     ANSWER 2 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN
     9001-92-7 REGISTRY
     Proteinase (9CI) (CA INDEX NAME)
OTHER NAMES:
      .alpha.-N-Benzoyl-DL-arginine-p-nitroanilide hydrolase
CN
CN
     537 Acidic protease
```

```
CN
     Actinase
      Alcalase 2.5LDX
CN
      Alkalase 2.4L FG
CN
     Alkalase 2.5L Type DX
CN
CN
      Alkalase 2.5L type X
CN
     Alkaline protease-L FG
CN
     ALP 901
CN
     Alphamalt BK 5020
CN
     Alphamalt LQ 4020
CN
     AO protease
     APL 901
CN
CN
     Aquatinase E
CN
     Arginine esterase
CN
     AS 1.398
     AS 10
CN
     Azocaseinase
CN
CN
     BAPAase
CN
     BAPNAase
CN
     Benzoyl arginine arylamidase
CN
     Benzoyl-DL-arginine-p-nitroanilide hydrolase
CN
     Bioprase 30L
CN
     Bioprase SP 4FG
CN
     Bioprotease A
     Bioprotease N 100P
CN
CN
     Biopurase
CN
     Biosoft PW
CN
     Carbonyl hydrolase
CN
     Casein endopeptidase
     Caseinase
CN
CN
     CL-5PG
CN
     Cleanase AP 100-PWC
CN
     Corolase 7089
CN
     Corolase L 10
CN
     DA 10
CN
     DA 10 (enzyme)
CN
     Denapsin 10P
CN
     Denatyme AP
CN
     Deozyme
CN
     Deterzyme L-600
     Durazyme 16.0L
CN
CN
     Endopeptidase
CN
     Endopeptidase 0
CN
     Endoprotease
CN
     Endoproteinase
CN
     Enzeco fungal acid protease
CN
     Enzylase K 40
CN
     Enzylon SAL
CN
     Enzylon SAL 300
ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for
     DISPLAY
     9001-93-8, 9012-23-1, 9040-76-0, 125498-72-8, 125752-86-5, 123779-18-0,
     124041-97-0, 120038-39-3, 120038-40-6, 105913-13-1, 118901-82-9,
     144906-30-9, 143404-30-2, 143404-41-5, 80804-52-0, 116267-38-0,
     117278-03-2, 117698-27-8, 118390-80-0
MF
     Unspecified
CI
     COM, MAN
LC
     STN Files:
                   ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
        CA, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN,
        CSCHEM, CSNB, DDFU, DIOGENES, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MSDS-OHS, NAPRALERT, NIOSHTIC, PIRA, PLASPEC*, PROMT, RTECS*,
        TOXCENTER, TULSA, USPAT2, USPATFULL, VTB
          (*File contains numerically searchable property data)
                      EINECS**, TSCA**
     Other Sources:
          (**Enter CHEMLIST File for up-to-date regulatory information)
DT.CA
       CAplus document type: Book; Conference; Dissertation; Journal; Patent;
       Report
       Roles from patents: ANST (Analytical study); BIOL (Biological study); CMBI (Combinatorial study); FORM (Formation, nonpreparative); MSC
        (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process);
        PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role
RLD.P Roles for non-specific derivatives from patents: ANST (Analytical study); BIOL (Biological study); MSC (Miscellaneous); OCCU (Occurrence);
        PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or
        reagent); USES (Uses)
RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological
```

```
study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU
        (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
        (Reactant or reagent); USES (Uses); NORL (No role in record)
RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical
        study); BIOL (Biological study); FORM (Formation, nonpreparative); MSC
        (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process);
        PRP (Properties); RACT (Reactant or reagent); USES (Uses)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
            40754 REFERENCES IN FILE CA (1907 TO DATE)
              491 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            40811 REFERENCES IN FILE CAPLUS (1907 TO DATE)
L5
     ANSWER 3 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN
     7784-30-7 REGISTRY
     Phosphoric acid, aluminum salt (1:1) (8CI, 9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Aluminum phosphate (Al(PO4)) (7CI)
OTHER NAMES:
     ALPO
CN
CN
     AlPO 11
     Alpo 5
CN
CN
     aluminophosphate (AlPO4)
CN
     Aluminum monophosphate
CN
     Aluminum orthophosphate
     Aluminum phosphate
CN
CN
     Aluminum phosphate (1:1)
     Aluphos
CN
CN
     Fabutit 320
     Fabutit 748
CN
CN
     FB 67
CN
     FFB 32
CN
     Fosfalugel
     Fosfalumina
CN
CN
     K-Bond 90
CN
     Monoaluminum phosphate
CN
     Phosphaljel
     Phosphalugel
CN
CN
     Phosphalujel
CN
     Phosphalutab
CN
     Phosphaluvet
CN
     Ulcocid
CN
     VPT
AR
     98499-64-0
     13765-93-0, 8022-59-1, 135151-77-8, 51668-55-4, 36201-72-6, 37324-42-8,
     93237-81-1, 89686-54-4, 52350-11-5
MF
     Al . H3 O4 P
CI
     COM
LC
     STN Files:
                  ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
       CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS,
       CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, DDFU, DETHERM*, DIPPR*, DRUGU,
       EMBASE, ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT, ENCOMPPAT2, GMELIN*, IFICDB,
       IFIPAT, IFIUDB, IMSCOSEARCH, IPA, MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC,
       PDLCOM*, PIRA, PROMT, RTECS*, TOXCENTER, TULSA, ULIDAT, USAN, USPAT2,
       USPATFULL, VETU, VTB
         (*File contains numerically searchable property data)
                     DSL**, EINECS**, TSCA**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
DT.CA Caplus document type: Book; Conference; Dissertation; Journal; Patent;
       Preprint; Report
RI. P
       Roles from patents: ANST (Analytical study); BIOL (Biological study);
       FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU
       (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
(Reactant or reagent); USES (Uses); NORL (No role in record)
RLD.P Roles for non-specific derivatives from patents: BIOL (Biological
       study); PREP (Preparation); PROC (Process); PRP (Properties); RACT
       (Reactant or reagent); USES (Uses)
RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological
       study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU
       (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
       (Reactant or reagent); USES (Uses); NORL (No role in record)
RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU
```

```
(Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)
CRN (7664-38-2)
```

```
Al
            6359 REFERENCES IN FILE CA (1907 TO DATE)
             108 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            6374 REFERENCES IN FILE CAPLUS (1907 TO DATE)
               4 REFERENCES IN FILE CAOLD (PRIOR TO 1967)
    ANSWER 4 OF 4 REGISTRY COPYRIGHT 2004 ACS on STN
    151-21-3 REGISTRY
RN
    Sulfuric acid monododecyl ester sodium salt (8CI, 9CI) (CA INDEX NAME)
CN
OTHER NAMES:
CN
    Adeka Hope LS 35
    Adeka Hope LS 90
CN
CN
    Akyposal NLS
CN.
    Akyposal SDS
CN
    Alscoap LN 40A
    Alscoap LN 90
CN
     Alscoap MP 90N
CN
CN
     Alscoap SP 40
CN
    Aquarex Me
     Avirol 101
CN
     Avirol SL 2010
CN
CN
     Berol 452
    Bio-Soft SDBS 60
CN
    Calfoam SLS 30
CN
     Carsonol SLS-S
CN
CN
    Conco Sulfate WAS
CN
     Cycloryl 21LS
    Cycloryl 580
CN
CN
    Dehydag Sulfate GL
CN
     Dermacide
CN
     Dodecyl sodium sulfate
CN
     Dodecyl sulfate sodium salt
CN
     Dreft
CN
    Duponol C
CN
     Duponol ME
CN
     Duponol QC
     Duponol WA
CN
CN
     Duponol WA Dry
CN
     Duponol WAQ
CN
     Duponol WAQE
CN
    Duponol WAQM
     Emal 10
CN
CN
     Emal 10 Needle
CN
     Emal 10 Powder
CN
     Emal 2F
     Emal 2F Needle
CN
CN
     Emal 2F30
CN
     Emal O
CN
     Emal OS
    Empicol 0303
Empicol 0303VA
CN
CN
CN
     Empicol BSD 70
CN
     Empicol LPZ
     Empicol LS 30
CN
CN
     Empicol LX 28
CN
     Empicol LX 28R
CN
     Empicol LX 42
CN
     Empicol LXSV 938U
     Empicol LXV
CN
     Empicol LY 28S
CN
ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for
```

```
DISPLAY
     12738-53-3, 12765-21-8, 8012-56-4, 1334-67-4, 1335-72-4, 172826-72-1,
DR
     121481-64-9, 58640-35-0, 57176-54-2, 64441-33-4, 129203-37-8, 51222-39-0,
     61711-39-5, 111726-87-5, 74433-77-5, 145269-44-9, 152155-52-7,
     156108-01-9, 191490-40-1, 237743-45-2, 303179-49-9
     C12 H26 O4 S . Na
     COM
CI
                   ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*,
T.C
     STN Files:
       BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT,
       CBNB, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DETHERM*, DIOGENES, DRUGU, EMBASE, ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT,
       ENCOMPPAT2, GMELIN*, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC, PDLCOM*, PHAR, PIRA, PROMT, RTECS*, TOXCENTER, TULSA, ULIDAT, USAN, USPAT2, USPATFULL, VETU, VTB
     (*File contains numerically searchable property data)
Other Sources: DSL**, EINECS**, TSCA**
          (**Enter CHEMLIST File for up-to-date regulatory information)
       CAplus document type: Book; Conference; Dissertation; Journal; Patent;
        Preprint; Report
       Roles from patents: ANST (Analytical study); BIOL (Biological study); MSC (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC
RL.P
        (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses);
       NORL (No role in record)
RLD.P Roles for non-specific derivatives from patents: ANST (Analytical
        study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP
        (Properties); USES (Uses)
RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological
        study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU
        (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
        (Reactant or reagent); USES (Uses); NORL (No role in record)
RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical
        study); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU
        (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
        (Reactant or reagent); USES (Uses)
CRN
     (151-41-7)
HO_3SO-(CH_2)_{11}-Me
       Na
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
            30993 REFERENCES IN FILE CA (1907 TO DATE)
              324 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            31054 REFERENCES IN FILE CAPLUS (1907 TO DATE)
               32 REFERENCES IN FILE CAOLD (PRIOR TO 1967)
FILE 'WPIX' ENTERED AT 11:27:38 ON 16 DEC 2004
COPYRIGHT (C) 2004 THE THOMSON CORPORATION
                               13 DEC 2004
FILE LAST UPDATED:
                                                   <20041213/UP>
MOST RECENT DERWENT UPDATE:
                                    200480
                                                    <200480/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE
>>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE,
    PLEASE VISIT:
 http://www.stn-international.de/training_center/patents/stn_guide.pdf <<<
>>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES. SEE
    http://thomsonderwent.com/coverage/latestupdates/
                                                                        <<<
```

FOR FURTHER DETAILS: http://www.thomsonderwent.com/dwpifv <<<

Search done by Noble Jarrell

DOCUMENTATION NOW AVAILABLE IN DERWENT WORLD PATENTS INDEX

<<<

>>> FOR INFORMATION ON ALL DERWENT WORLD PATENTS INDEX USER

http://thomsonderwent.com/support/userguides/

>>> NEW! FAST-ALERTING ACCESS TO NEWLY-PUBLISHED PATENT

GUIDES, PLEASE VISIT:

FIRST VIEW - FILE WPIFV.

```
>>> NEW DISPLAY FORMAT HITSTR ADDED ALLOWING DISPLAY OF HIT STRUCTURES WITHIN THE BIBLIOGRAPHIC DOCUMENT <
```

>>> SMILES and ISOSMILES strings are no longer available as
Derwent Chemistry Resource display fields <<<</pre>

```
=> d all 18 tot
```

```
L8 ANSWER 1 OF 1 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN AN 2001-168447 [17] WPIX
```

DNC C2001-050284

TI Novel multivalent immunogenic composition for conferring protection against infection caused by Hameophilus influenzae and Moraxella catarrhalis comprises four antigens derived from each of the two microorganisms.

DC B04 D16

IN KLEIN, M H; LOOSMORE, S M; SASAKI, K; YANG, Y

PA (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD

CYC 95

PI WO 2001005424 A2 20010125 (200117) \* EN 58 A61K039-00 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2000059586 A 20010205 (200128) A61K039-00 EP 1200122 A2 20020502 (200236) EN A61K039-116

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

US 6391313 B1 20020521 (200239) A61K039-116 <-AU 767096 B 20031030 (200382) A61K039-00
NZ 516819 A 20031219 (200404) A61K039-00

ADT WO 2001005424 A2 WO 2000-CA811 20000711; AU 2000059586 A AU 2000-59586 20000711; EP 1200122 A2 EP 2000-945494 20000711, WO 2000-CA811 20000711; US 6391313 B1 US 1999-353617 19990715; AU 767096 B AU 2000-59586 20000711; NZ 516819 A NZ 2000-516819 20000711, WO

2000-CA811 20000711 FDT AU 2000059586 A Based on WO 2001005424; EP 1200122 A2 Based on WO 2001005424; AU 767096 B Previous Publ. AU 2000059586, Based on WO 2001005424; NZ 516819 A Based on WO 2001005424

PRAI US 1999-353617 19990715

IC ICM A61K039-00; A61K039-116

ICS A61P031-04

AB WO 200105424 A UPAB: 20010328

NOVELTY - A multivalent immunogenic composition (I) for conferring protection in a host against disease caused by both Hameophilus influenzae (HI) and Moraxella catarrhalis (MC) comprising four different antigens, of which at least one antigen is from HI and one antigen is from MC, is new. Additionally three of the antigens of (I) are adhesins, and one is from MC

ACTIVITY - Auditory; antibacterial.

MECHANISM OF ACTION - Vaccine.

Groups of five BALB/C mice were immunized subcutaneously on days 1,29 and 43 with one of the mouse H91A Hin47 + rHMW + rHia + r200 kDa vaccines. Blood samples were taken on days 0, 14, 28, 42 and 56. Groups of five Hartley outbreed guinea pigs were immunized intramuscularly on days 1, 29 and 43 with the vaccine as described above. Blood samples were taken on days 0, 14, 28, 42 and 56. Anti-H91A Hin47, anti-rHMW, anti-rHia and anti-r200 kDa IgG antibody titers were determined by antigen specific enzyme linked immunosorbant assays (ELISAs). The results of the immunogenicity studies showed that the final bleed sera obtained from mice immunized with 0.3 mu g, or 3.0 mu g each of H91A Hin47 + rHMW + rHia with 0, 0.3, 1.0, 3.0 or 10.0 mu g of added r200 kDa, all had high antibody titers to H91A Hin47 component. The final bleed sera obtained from the mice immunized with 3.0 mu g each of H91A Hin47 + rHMW + rHia with 0, 0.3, 1.0, 3.0 or 10.0 mu g of added r200 kDa, all had high titer antibodies to the rHMW apparent enhancing or inhibiting effect on the anti-rHMW response with the addition of the  $r200\ kDa$  component. Mice immunized with 0.3 mu g each of H91A Hin 47 + HMW + rHia with 0, 0.3, 1.0, 3.0 or 10.0 mu g of added r200 kDa, all had high titer antibodies to the rHia component. There was no apparent enhancing or inhibiting effect on the anti-rHia response with the addition of the r200 kDa component. The final bleed sera obtained from guinea pigs immunized with 25 mu g or 50 mu g each of H91A Hin47 + rHMW + rHia with 0, 25, 50 or 100 mu g of added r200 kDa, all had high

RMIN

titer antibodies to the H91A Hin47 component. Also final bleed sera obtained from guinea pigs immunized with 25 mu g or 50 mu g each of H91A Hin47 + rHMW + rHia with 0, 25, 50 or 100 mu g of added r200 kDa, all had titer antibodies to the rHMW component. There was no apparent enhancing or inhibiting effect on the anti-rHMW response upon the addition of the r200 kDa antigen.

USE - (I) is useful for immunizing a host against infection caused by both HI and MC including otitis media (claimed).

ADVANTAGE - The multivalent vaccine can confer protection against encapsulated and unencapsulated HI and MC diseased in a safe and efficient manner.

Dwg.0/14

FS CPI

FA AB; DCN

MC CPI: B04-B04C1; B14-A01; B14-A01A; B14-N02; B14-S11B; D05-C02; D05-H07; D05-H12F

=> b home

FILE 'HOME' ENTERED AT 11:27:45 ON 16 DEC 2004

= :

```
=> d his
```

```
(FILE 'HOME' ENTERED AT 11:24:56 ON 16 DEC 2004)
     FILE 'HCAPLUS' ENTERED AT 11:25:27 ON 16 DEC 2004
                E WO2000-CA811/AP, PRN
              1 WO2000-CA811/AP, PRN
Ll
              1 US6391313/PN
L2
L3
              1 L1-2
     FILE 'REGISTRY' ENTERED AT 11:26:35 ON 16 DEC 2004
     FILE 'HCAPLUS' ENTERED AT 11:26:37 ON 16 DEC 2004
                TRA L3 1- RN :
                                      4 TERMS
L4
     FILE 'REGISTRY' ENTERED AT 11:26:37 ON 16 DEC 2004
L5
              4 SEA L4
     FILE 'WPIX' ENTERED AT 11:26:42 ON 16 DEC 2004
                E WO2000-CA811/AP, PRN
              1 WO2000-CA811/AP, PRN
L6
L7
              1 US6391313/PN
L8
              1 L6-7
     FILE 'HCAPLUS' ENTERED AT 11:46:19 ON 16 DEC 2004
                E IMMUNOSTIMULANTS/CT
                E E3+ALL
L9
          16542 IMMUNOSTIMULANTS+NT/CT
                E IMMUNE ADJUVANTS/CT
                E E3+ALL
L10
           9606 IMMUNE ADJUVANTS/CT OR L9 (L) ADJUV?
                E VACCINES/CT
                E E3+ALL
                E CONTRACEPTIVES/CT
                E E3+ALL
            564 CONTRACEPTIVES/CT (L) ?IMMUN?/BI
L11
          40848 VACCINES/CT
1.12
                E IMMUNITY/CT
                E E3+ALL
          63934 IMMUNITY+NT/CT
L13
                E IMMUNOTHERAPY/CT
                E E3+ALL
L14
          12699 IMMUNOTHERAPY+NT/CT
                E THERAPEUTICS/CT
                E E3+ALL
                E E2
                E E3+ALL
          29488 THERAPY+OLD, NT/CT (L) ?IMMUN?/BI
L15
                E RADIOTHERAPY/CT
                E E3+ALL
                E HAEMOPHILUS INFLUENZAE/CT
                E E3+ALL
                E E4+ALL
L16
           7294 HAEMOPHILUS+OLD, NT/CT
                E MORAXELLA CATARRHALIS/CT
                E E3+ALL
                E E4
                E E3+ALL
           1861 MORAXELLA+OLD, NT/CT
L17
                E ANTIGENS/CT
                E E3+ALL
                QUE ANTIGENS+NT/CT
L18
             80 L16 AND L17 AND L18
L19
                E ADHESIN/CT
                E E5+ALL
                E AGGLUTININS AND LECTINS/CT
                E E3+ALL
1.20
           2116 "AGGLUTININS AND LECTINS"+OLD, NT/CT (L) ?ADHESIN?/BI
L21
              2 L19 AND L20
                E LOOSMORE S/AU
L22
             83 E3-7
                E YANG Y/AU
L23
            786 E3,E26
                E YANG YAN/AU
L24
            321 E3,E31
                E YANG, YANPING/AU
```

Graser 10/030313

Page 2

```
1 Lord
                E KLEIN ,/AU
                E KLEIN M/AU
L25
            513 E3,E12
                E KLEIN MICHAEL/AU
L26
            140 E3,E12
                E SASAKI K/AU
            702 E3-6
L27
                E SASAKI KEN/AU
L28 ·
            889 E3-10
L29
            182 (AVENTIS (1A) PASTEUR)/CS, PA
              3 L19 AND L22-29
L30
L31
             77 L19 NOT L30
L32
             57 L31 AND L9-15
                QUE PY<=1999 OR AY<=1999 OR PRY<=1999 OR PD<19990715 OR PRD<199
L33
             26 L32 AND L33
L34
                SEL AN 4 15 23 24 L34
L35
             22 L34 NOT E1-8
              1 L20 AND L35
L36
             21 L35 NOT L36
L37
                E OTITIS MEDIA/CT
                E E3+ALL
                E EAR, DISEASE/CT
                E E3=ALL
                E EAR, DISEASE/CT
                E E3+ALL
L38
           2966 "EAR, DISEASE"+NT/CT
                E EAR/CT
                E E3=ALL
                E EAR/CT
                E E3+ALL
L39
           1839 EAR+OLD, NT/CT (L) DISEAS?
           9750 EAR+OLD, NT/CT
L40
L41
            818 L38-40 (L) OTITIS(1A) MEDIA
L42
              3 L35 AND L41
L43
              4 L36 OR L42
             18 L35 NOT L43
L44
     FILE 'WPIX' ENTERED AT 12:43:52 ON 16 DEC 2004
L45
          65479 A61K039/IPC OR (B04-B04C? OR C04-B04C? OR B04-G01 OR C04-G01)/M
          28924 (B12-A01 OR C12-A01 OR B14-A01A OR C14-A01A)/MC OR MORAXELLA/BI
L46
          28459 (B02-V02 OR C02-V02 OR B14-S11? OR C14-S11? OR D05-H07)/MC OR (
L47
L48
            989 L45 AND L46 AND L47
            205 (AVENTIS (1A) PASTEUR)/CS,PA
L49
                E LOOSMORE S/AU
             39 E3-4
L50
                E YANG Y/AU
L51
           1938 E3, E19
                E KELIN M/AU
                E KLEIN M/AU
L52
            333 E3.E13
                E SASAKI K/AU
           1213 E3-7
L53
L54
             29_L48 AND L49-53
            960 L48 NOT L54
L55
              4 L55 AND ADHESIN?/BIX
L56
L57
           8186 (B14-N02 OR C14-N02 OR B12-L04 OR C12-L04)/MC
L58
             40 L55 AND L57
              0 L58 AND ADHESIN?/BIX
L59
             18 L58 NOT (PY>1999 OR PRY>1999 OR AY>1999)
1.60
                SEL AN 1-3 10 12
              5 E1-5 AND L60
=> b hcap
FILE 'HCAPLUS' ENTERED AT 13:09:10 ON 16 DEC 2004
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
```

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 16 Dec 2004 VOL 141 ISS 25 FILE LAST UPDATED: 15 Dec 2004 (20041215/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

```
=> d all 144 tot
    ANSWER 1 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
    2002:505235 HCAPLUS
AN
DN
    137:62165
    Entered STN: 05 Jul 2002
ED
    Producing antibodies with attenuated bacteria with altered DNA adenine
TI
    methylase activity
    Mahan, Michael J.; Heithoff, Douglas M.; Low, David A.; Sinsheimer, Robert
IN
    L.
PA
    USA
    U.S. Pat. Appl. Publ., 44 pp., Cont.-in-part of U.S. Ser. No. 612,116.
    CODEN: USXXCO
рΤ
    Patent
    English
LΑ
IC
    ICM A61K039-02
    ICS C12N001-21
NCL
    424200100
CC
    15-2 (Immunochemistry)
     Section cross-reference(s): 3, 14
FAN.CNT 7
    PATENT NO.
                         KIND
                                DATE
                                            APPLICATION NO.
    US 2002086032
                               20020704
                         A1
                                           US 2001-927896
                                                                   20010809 <
PRAI US 1999-183043P
                          Р
                                19990202 <--
                         р
                                          <--
```

US 1999-198250P 19990505 US 2000-495614 A2 20000201 US 2000-612116 A2 20000707 CLASS

PATENT NO. CLASS PATENT FAMILY CLASSIFICATION CODES US 2002086032 TCM A61K039-02

ICS C12N001-21 NCL 424200100

A61K039/002; A61K039/02; A61K039/106; A61K039/112 US 2002086032 ECLA The present invention is directed towards methods of inducing antibodies using an attenuated strain of pathogenic bacteria (e.g., Haemophilus, Escherichia coli, and/or Salmonella) having non-reverting genetic mutations relative to the wild-type organism which alter activity of DNA adenine methylase (Dam). The invention further includes compns. comprised of the attenuated bacteria and methods using these compns. to elicit an immune response and immunize a subject with highly specific antibodies. The invention also provides methods producing antibodies to heterologous antiqens which the attenuated bacteria are engineered to produce.

antibody bacteria DNA adenine methylase; vaccine Salmonella vector DNA ST adenine methylase

IT Vaccines

(AIDS; attenuated bacteria with altered DNA adenine methylase activity for expression of heterologous antigens of HIV)

IT Antibodies and Immunoglobulins

RL: BSU (Biological study, unclassified); BIOL (Biological study) (IGG; attenuated bacteria with altered DNA adenine methylase activity for induction of)

IT Animal virus Arbovirus

Ascaris lumbricoides Aspergillus fumigatus Astrovirus Bacillus anthracis Blastomyces dermatitidis Bordetella pertussis Borrelia burgdorferi Campylobacter Candida Chlamydia pneumoniae Chlamydia trachomatis

Clostridium tetani Coccidioides Cryptococcus neoformans Cytomegalovirus Dengue virus Entamoeba histolytica Giardia lamblia Helicobacter pylori Hepatitis A virus Hepatitis B virus Hepatitis C virus Hepatitis E virus Hepatitis GB virus C/G Hepatitis delta virus Histoplasma capsulatum Human coxsackievirus Human echovirus Human herpesvirus 2 Human herpesvirus 3 Human herpesvirus 4 Human papillomavirus Human parainfluenza virus Human poliovirus Influenza virus Japanese encephalitis virus Leptospira Measles virus Moraxella catarrhalis Mycobacterium leprae Mycobacterium tuberculosis Mycoplasma pneumoniae Neisseria gonorrhoeae Neisseria meningitidis Norwalk virus Paracoccidioides brasiliensis Paramyxovirus Parasite Pinworm Plasmodium (malarial genus) Pseudomonas aeruginosa Rabies virus Respiratory syncytial virus Rhinovirus Rotavirus Rubella virus Schistosoma Staphylococcus aureus Staphylococcus saprophyticus Streptococcus group A Streptococcus group B Taenia Toxoplasma gondii Treponema pallidum Trichomonas vaginalis Variola virus (attenuated bacteria with altered DNA adenine methylase activity for expression of heterologous antigens of) Mycosis (attenuated bacteria with altered DNA adenine methylase activity for expression of heterologous antigens of fungi associated with) Sexually transmitted diseases (attenuated bacteria with altered DNA adenine methylase activity for expression of heterologous antigens of microorganisms associated with) Gene, microbial RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (dam; of attenuated bacteria with altered DNA adenine methylase activity) Antigens RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (heterologous; expression in attenuated bacteria with altered DNA adenine methylase activity) Escherichia Eubacteria Haemophilus Salmonella Vibrio

IT

IT

IT

IT

IT

```
Yersinia
        (immunostimulation by attenuated bacteria with altered DNA adenine
        methylase activity)
     Vaccines
IT
        (of attenuated bacteria with altered DNA adenine methylase activity)
TT
     Antigens
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (tumor-associated; expression in attenuated bacteria with altered DNA
        adenine methylase activity)
IT
     Bos taurus
     Gallus domesticus
     Human
        (vaccination with attenuated bacteria expressing altered DNA adenine
        methylase activity and heterologous antigens)
IT
     Food poisoning
        (vaccination with attenuated bacteria expressing altered DNA adenine
        methylase activity and heterologous antigens in relation to)
IT
     Anti-AIDS agents
        (vaccines; attenuated bacteria with altered DNA adenine methylase
        activity for expression of heterologous antigens of HIV)
IT
     69553-52-2, Dam methylase
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (immunostimulation by attenuated bacteria with altered DNA adenine
        methylase activity)
    ANSWER 2 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
L44
AN
     2002:488067 HCAPLUS
DN
     137:62150
     Entered STN: 28 Jun 2002
ED
TI
     Bacteria with mutated DNA adenine methylase for use as vaccine and
     screening or development of antimicrobial or antibacterial agents
    Mahan, Michael J.; Heithoff, Douglas M.; Low, David A.; Sinsheimer, Robert
IN
PA
     USA
     U.S. Pat. Appl. Publ., 44 pp., Cont.-in-part of U.S. Ser. No. 612,116.
so
     CODEN: USXXCO
DT
     Patent
LΑ
     English
IC
     ICM A61K039-02
         C12N001-21; C12N015-74
     ICS
NCL
     424200100
     15-2 (Immunochemistry)
     Section cross-reference(s): 1, 3, 10, 17
FAN.CNT 7
                                            APPLICATION NO.
     PATENT NO.
                         KTND
                                DATE
                                                                    DATE
                         ----
                                -----
     US 2002081317
                                20020627
                                            US 2001-927788
                                                                    20010809
                          Al
     ZA 2001005305
                          Α
                                20020627
                                             ZA 2001-5305
                                                                    20010627
                                19990202
PRAI US 1999-183043P
                          Р
     US 1999-241951
                          Α
                                19990202
     US 1999-198250P
                          P
                                19990505
     US 1999-305603
                          A
                                19990505
     US 2000-495614
                                20000201
                          A2
    US 2000-612116
                                20000707
                          Α2
CLASS
 PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
US 2002081317
                 ICM
                        A61K039-02
                        C12N001-21; C12N015-74
                 ICS
                 NCL
                        424200100
                        A61K039/002; A61K039/02; A61K039/106; A61K039/112
US 2002081317
                ECLA
    Immunogenic compns. are disclosed which are comprised of bacteria which
AB
     are pathogenic in their native state but which are rendered non-pathogenic
     in a manner which alters the native level or activity of DNA adenine
     methylase (dam). The genome is also artificially engineered to express a
     heterologous antigen such as an immunogenic antigen of a virus, protozoa,
     parasite or fungi. The microorganism with mutated dam is also useful for
     identifying or developing antimicrobial or antibacterial agents.
     DNA adenine methylase mutation microorganism vaccine; antimicrobial
ST
     antibacterial vaccine antigen DNA adenine methylase mutation
IT
    Hepatitis
        (A; bacteria with mutated DNA adenine methylase for use as vaccine and
```

Hepatitis
(B; bacteria with mutated DNA adenine methylase for use as vaccine and

screening or development of antimicrobial and antibacterial agents)

IT

Graser 10/030313

Page 6

screening or development of antimicrobial and antibacterial agents) IT Hepatitis (C; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) IT (D; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) TT Hepatitis (E; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) Antibodies and Immunoglobulins IT RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (IgG; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) IT Haemophilus influenzae (NT; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) IT Toxins RL: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (Shiga; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) TΤ Drug screening (antibacterial; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) IT Fungi Parasite Protozoa Virus (antigen; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) TТ Arbovirus Ascaris lumbricoides Aspergillus fumigatus Astrovirus Bacillus anthracis Blastomyces dermatitidis Bordetella pertussis Borrelia burgdorferi Bos taurus Campylobacter Candida Chlamydia pneumoniae Chlamydia trachomatis Coccidioides immitis Cryptococcus neoformans Cytomegalovirus Dengue virus Drug delivery systems Entamoeba histolytica Eubacteria Food poisoning Gallus domesticus Genetic vectors Giardia lamblia Helicobacter pylori Hepatitis GB virus C/G Hepatitis virus Herpesviridae Histoplasma capsulatum Human Human coxsackievirus Human echovirus Human herpesvirus Human herpesvirus 1 Human herpesvirus 2 Human herpesvirus 3 Human herpesvirus 4 Human immunodeficiency virus Human papillomavirus Human parainfluenza virus Human poliovirus Influenza virus

```
Japanese encephalitis virus
Leptospira
Mammalia
Measles virus
Molecular cloning
   Moraxella catarrhalis
Mutagenesis
Mycobacterium leprae
Mycobacterium tuberculosis
Mycoplasma pneumoniae
Mycosis
Neisseria gonorrhoeae
Neisseria meningitidis
Norwalk virus
Paracoccidioides brasiliensis
Paramyxovirus
Pathogen
Pathogenic bacteria
Pinworm
Plasmodium (malarial genus)
Pseudomonas aeruginosa
Rabies virus
Respiratory syncytial virus
Rhinovirus
Rodentia
Rotavirus
Rubella virus
Salmonella
Salmonella enteritidis
Salmonella typhi
Salmonella typhimurium
Schistosoma
Sexually transmitted diseases
Shigella
Staphylococcus saprophyticus
Streptococcus group A
Streptococcus group B
Taenia
Tetanus
Toxoplasma gondii
Treponema pallidum
Trichomonas vaginalis
Typhoid fever
  Vaccines
 Vibrio
Vibrio cholerae
Yersinia
Yersinia pseudotuberculosis
    (bacteria with mutated DNA adenine methylase for use as vaccine and
    screening or development of antimicrobial and antibacterial agents)
Antibodies and Immunoglobulins
  Antigens
RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
(Uses)
    (bacteria with mutated DNA adenine methylase for use as vaccine and
    screening or development of antimicrobial and antibacterial agents)
Gene, microbial
RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
    (bacteria with mutated DNA adenine methylase for use as vaccine and
    screening or development of antimicrobial and antibacterial agents)
    (bacterial; bacteria with mutated DNA adenine methylase for use as
    vaccine and screening or development of antimicrobial and antibacterial
    agents)
Antibacterial agents
Antimicrobial agents
    (development; bacteria with mutated DNA adenine methylase for use as
    vaccine and screening or development of antimicrobial and antibacterial
   agents)
Immunity
    (disorder, antigen; bacteria with mutated DNA adenine methylase for use
    as vaccine and screening or development of antimicrobial and
    antibacterial agents)
Escherichia coli
```

TΤ

IT

Page 8

Graser 10/030313 (enterotoxigenic; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) Respiratory tract, disease IT Urinary tract, disease (infection; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) IT Intestinal bacteria (pathogenic, infection; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents) Antigens

IT

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(tumor-associated; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents)

IT Haemophilus influenzae

(type b; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents)

IT

(uropathogenic; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents)

IT Infection

> (variola; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents)

IT Infection

(vector-born; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents)

IT Infection

> (viral; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents)

TT 69553-52-2. Dam methylase

RL: BSU (Biological study, unclassified); REM (Removal or disposal); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (mutation; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial and antibacterial agents)

IT 439624-92-7 439624-93-8 439624-94-9 439624-95-0

RL: PRP (Properties)

(unclaimed sequence; bacteria with mutated DNA adenine methylase for use as vaccine and screening or development of antimicrobial or antibacterial agents)

- L44 ANSWER 3 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
- 2002:107146 HCAPLUS AN
- DN 136:166052
- Entered STN: 10 Feb 2002
- TI Vaccine composition
- Berthet, François-Xavier Jacques; Dalemans, Wilfried; Denoel, Philippe; Dequesne, Guy; Feron, Christiane; Garcon, Nathalie; Lobet, Yves; Poolman, IN Jan; Thiry, Georges; Thonnard, Joelle; Voet, Pierre
- PA Smithkline Beecham Biologicals SA, Belg.
- PCT Int. Appl., 125 pp. so
- CODEN: PIXXD2
- DT Patent
- English LΑ
- IC ICM A61K039-00
- 15-2 (Immunochemistry) CC

Section cross-reference(s): 3

FAN.CNT 2

	PATENT	KIN	כ	DATE			APPLICATION NO.						DATE						
						-													
PΙ	WO 2002009746				A2		20020207			WO 2001-EP8857						20010731			
	WO 2002	A3 20020613																	
	WO 2002	C1		2002	1114														
	W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,		
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,		
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	KZ,	LC,	LK,	LR,		
		LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PL,	PT,		
		RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	TZ,	UA,	ŪĠ,	US,		
		UΖ,	VN,	YU,	ZA,	ZW,	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM				

```
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                 20020529
                                             EP' 2000-956369
                                                                      20000731 <--
     EP 1208214
                          A2
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL
                          AA
                                 20020207
                                              CA 2001-2425037
     CA 2425037
                                                                      20010731
                                              AU 2001-85856
     AU 2001085856
                          A5
                                 20020213
                                                                      20010731
     EP 1307224
                          A2
                                 20030507
                                             EP 2001-965152
                                                                      20010731
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
     US 2004126389
                                 20040701
                                              US 2003-343561
                                                                      20030915
                          Α1
PRAI EP 2000-956369
                          Α
                                 20000731
     GB 2001-3170
                          Α
                                 20010208
     GB 1999-18319
                          Α
                                 19990803
                                           <--
     WO 2000-EP7424
                          W
                                 20000731
     WO 2001-EP8857
                          W
                                 20010731
CLASS
                 CLASS PATENT FAMILY CLASSIFICATION CODES
PATENT NO.
WO 2002009746
                 ICM
                        A61K039-00
US 2004126389 ECLA
                       A61K039/02; A61K039/39
    The present invention relates to the field of vaccine formulation,
     particularly the field of novel adjuvant compns. comprising outer membrane
     vesicles (or blebs), and advantageous methods of detoxifying these
     compns., and advantageous methods of use of such adjuvants. The novel
     adjuvant for Gram-neg. bacterial vaccine is a capsular polysaccharide or
     detoxified lipid A portion of LPS derived from engineered Neisseria
     meningitidis serogroup A, B, Y or W; Hemophilus influenzae; Streptococcus
     pneumoniae; or Moraxella catarrhalis. These engineered bacteria have
     reduced or switched off expression of one or more gene selected from htrB,
     msbB, .pxK, pmrA, pmrB, pmrE, pmrF, galE, siaA, siaB, siaC, siaD, ctrA,
     ctrB, ctrC and ctrD. Vaccines comprising the adjuvant and
     pathogen-derived antigen is especially useful for protecting elderly patients
     against the pathogen.
    vaccine adjuvant outer membrane vesicle bleb; Gram neg bacteria bleb prepn adjuvant; Neisseria meningitidis bleb detoxified lipid A; Streptococcus
ST
     pneumoniae Hemophilus influenzae capsular polysaccharide
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (D15; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
     Proteins
     RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
     THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
        (D; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
     Proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (OMP (outer membrane protein); outer membrane vesicles or detoxified
        lipid A as adjuvant for Gram-neg. bacterial vaccines)
IT
     Immunostimulants
        (adjuvants; outer membrane vesicles or detoxified lipid A as
        adjuvant for Gram-neg. bacterial vaccines)
IT
     Organelle
        (bleb; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
    Glycolipids
     Lipopolysaccharides
     Polysaccharides, biological studies
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (capsular; outer membrane vesicles or detoxified lipid A as adjuvant
        for Gram-neg. bacterial vaccines)
TT
    Drug delivery systems
        (carriers; outer membrane vesicles or detoxified lipid A as adjuvant
        for Gram-neg. bacterial vaccines)
IT
    Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (cps; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
TT
     Gene, microbial
```

Page 10

Graser 10/030313 RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process) (ctrA; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) IT Gene, microbial RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process) (ctrB; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) Gene, microbial RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process) (ctrC; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) TT Gene, microbial RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process) (ctrD; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) IT Aging, animal (elderly; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) IT Gene, microbial RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process) (galE; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) IT Proteins RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified); BIOL (Biological study); PREP (Preparation) (green fluorescent; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) Neisseria meningitidis IT (group A; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) IT Neisseria meningitidis (group B; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) Neisseria meningitidis (group W; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) IT Neisseria meningitidis (group Y; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) Gene. microbial RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process) (hsf; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) Gene, microbial RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process) (htrB; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines) Gene, microbial RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process) (lpxK; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines)

TT

IT

IT

TT

IT Gene, microbial

RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)

(msbB; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines)

IT Gene, microbial

RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)

(nspA; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines)

Gene, microbial

RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)

(omp85; outer membrane vesicles or detoxified lipid A as adjuvant for Gram-neg. bacterial vaccines)

IT DNA sequences

Detergents

Gram-negative bacteria
Haemophilus influenzae

```
Moraxella catarrhalis
     Neisseria meningitidis
     Streptococcus pneumoniae
       Vaccines
        (outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
     Promoter (genetic element)
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
     Antigens
     Polysaccharides, biological studies
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
TT
     Cell wall
        (outer membrane, vesicles; outer membrane vesicles or detoxified lipid
        A as adjuvant for Gram-neg. bacterial vaccines)
TΤ
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)
        (pilQ; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)
        (pldA; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neq. bacterial vaccines)
IT
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)
        (pmrA; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
TT
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (pmrB; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
     Gene, microbial
TT
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (pmrE; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (pmrF; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
     Infection
        (pneumococcal; outer membrane vesicles or detoxified lipid A as
        adjuvant for Gram-neg. bacterial vaccines)
     Gene, microbial
IT
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (porA; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (porB; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (siaA; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (siaB; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
TТ
     Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
```

```
(Biological study); PROC (Process)
        (siaC; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
    Gene, microbial
     RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL
     (Biological study); PROC (Process)
        (siaD; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
    Gene, microbial
    RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)
        (tbpA; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
    Haemophilus influenzae
        (type b; outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IT
    1404-24-6, Polymyxin A
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (chimeric; outer membrane vesicles or detoxified lipid A as adjuvant
        for Gram-neg. bacterial vaccines)
    397430-36-3, DNA (Synthetic plasmid vector CMK(+))
    RL: BSU (Biological study, unclassified); BUU (Biological use,
    unclassified); PRP (Properties); BIOL (Biological study); USES (Uses)
        (nucleotide sequence; outer membrane vesicles or detoxified lipid A as
        adjuvant for Gram-neg. bacterial vaccines)
                  397430-38-5
                                397430-39-6
                                              397430-40-9
IT
    397430-37-4
     397430-42-1
    RL: BSU (Biological study, unclassified); PRP (Properties); REM (Removal
     or disposal); BIOL (Biological study); PROC (Process)
        (nucleotide sequence; outer membrane vesicles or detoxified lipid A as
        adjuvant for Gram-neg. bacterial vaccines)
     83-44-3
IT
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (outer membrane vesicles or detoxified lipid A as adjuvant for
        Gram-neg. bacterial vaccines)
IΤ
    397431-26-4, 4: PN: WOO209746 SEQID: 4 unclaimed DNA
                                                            397431-27-5, 5: PN:
     WOO209746 SEQID: 5 unclaimed DNA 397431-28-6, 6: PN: WOO209746 SEQID: 6
    unclaimed DNA 397431-29-7, 8: PN: WO0209746 SEQID: 8 unclaimed DNA
     397431-30-0, 9: PN: WO0209746 SEQID: 9 unclaimed DNA
                                                            397431-31-1
     397431-32-2
                  397431-33-3
                                397431-34-4
                                               397431-35-5
                                                             397431-36-6
     397431-37-7
                   397431-38-8
                                 397431-39-9
                                               397431-40-2
                                                             397431-41-3
     397431-42-4
                   397431-43-5
                                 397431-44-6
                                               397431-45-7
                                                             397431-46-8
                   397431-48-0
                                 397431-49-1
                                               397431-50-4
                                                             397431-51-5
     397431-47-9
                   397431-53-7
                                 397431-54-8
                                               397431-55-9
                                                             397431-56-0
     397431-52-6
     397431-57-1
                   397431-58-2
                                 397431-59-3
                                               397431-60-6
                                                             397431-61-7
                                 397431-64-0
                                               397431-65-1
     397431-62-8
                  397431-63-9
                                                             397431-66-2
     397431-67-3
                   397431-68-4
                                 397431-69-5
                                               397431-70-8
                                                             397431-71-9
                   397431-73-1
                                 397431-74-2
                                               397431-75-3
                                                             397431-76-4
    397431-72-0
     397431-77-5
                   397431-78-6
                                 397431-79-7
                                               397431-80-0
                                                             397431-81-1
     397431-82-2
                   397431-83-3
                                 397431-84-4
                                               397431-85-5
                                                              397431-86-6
    397431-87-7
                  397431-88-8
                                 397431-89-9
                                               397431-90-2
                                                             397431-91-3
                   397431-93-5
                                 397431~94-6
                                               397431-95-7
                                                             397431-96-8
    397431-92-4
                                 397431-99-1
                   397431-98-0
                                               397432-00-7
                                                             397432-01-8
    397431-97-9
     397432-02-9
                   397432-03-0
                                 397432-04-1
                                               397432-05-2
                                                             397432-06-3
     397432-07-4
                   397432-08-5
                                 397432-09-6
                                               397432-10-9
                                                             397432-11-0
    397432-12-1
                  397432-13-2
                                 397432-14-3
                                               397432-15-4
                                                             397432-16-5
                                 397432-19-8
    397432-17-6
                   397432-18-7
                                               397432-20-1
                                                             397432-21-2
    397432-22-3
                   397432-23-4
                                 397432-24-5
                                               397432-25-6
                                                             397432-26-7
     397432-27-8
                   397432-28-9
                                 397432-29-0
                                               397432-30-3
                                                             397432-31-4
     397432-32-5
                   397432-33-6
                                 397432-34-7
                                               397432-35-8
                                                             397432-36-9
                                 397432-39-2
                                                             397432-41-6
    397432-37-0
                   397432-38-1
                                               397432-40-5
    397432-42-7
                   397432-43-8
                                 397432-44-9
                                               397432-45-0
                                                             397432-46-1
     397432-47-2
                   397432-48-3
                                 397432-49-4
                                               397432-50-7
                                                             397432-51-8
                                 397432-54-1
                                               397432-55-2
     397432-52-9
                   397432-53-0
                                                             397432-56-3
                   397432-58-5
                                 397432-59-6
     397432-57-4
                                               397432-60-9
                                                             397432-61-0
                   397432-63-2
                                 397432-64-3
                                               397432-65-4
                                                              397432-66-5
     397432-62-1
     397432-67-6
                   397432-68-7
                                 397432-69-8
                                               397432-70-1
                                                             397432-71-2
                                 397432-74-5
     397432-72-3
                   397432-73-4
    RL: PRP (Properties)
        (unclaimed nucleotide sequence; vaccine composition)
L44 ANSWER 4 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
AN
    2000:688113 HCAPLUS
DN
    133:265640
```

```
Entered STN: 29 Sep 2000
ED
ΤI
     Bacterial polysaccharide antigen vaccine
     Capiau, Carine; Deschamps, Marguerite; Desmons, Pierre Michel; Laferriere,
TN
     Craig Antony Joseph; Poolman, Jan; Prieels, Jean-paul
PA
     Smithkline Beecham Biologicals SA, Belg.
so
     PCT Int. Appl., 79 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
IC
     ICM A61K039-385
     ICS A61K039-39; A61K039-02; A61K039-005; A61K039-116; A61P031-04
CC
     15-2 (Immunochemistry)
     Section cross-reference(s): 3
     PATENT NO.
                         KIND
                                DATE
                                             APPLICATION-NO.
                                                                    DATE
                         ----
PΤ
     WO 2000056360
                          A2
                                20000928
                                             WO 2000-EP2468
                                                                    20000317 <--
                                20010125
     WO 2000056360
                          А3
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
             CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
             ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
             SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA,
             ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
                        FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
             DK. ES. FI.
             CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     CA 2366314
                          AA
                                20000928
                                             CA 2000-2366314
                                                                    20000317 <--
                                             NZ 2000-513841
     NZ 513841
                          Α
                                 20010928
                                                                    20000317 <--
     EP 1163000
                          A2
                                 20011219
                                             EP 2000-912626
                                                                    20000317 <--
         R:
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
                                             TR 2001-200102739
     TR 200102739
                          T2
                                20011221
                                                                    20000317 <--
     BR 2000009163
                          Α
                                 20011226
                                             BR 2000-9163
                                                                    20000317 <--
     TR 200102735
                          T2
                                 20020422
                                             TR 2001-200102735
                                                                    20000317 <---
     TR 200102736
                          T2
                                 20020422
                                             TR 2001-200102736
                                                                    20000317 <--
                                             AU 2000-34307
                                                                    20000317 <--
     AII 750913
                                20020801
                          B2
     AU 2000034307
                          Α5
                                20001009
     JP 2002540075
                          Т2
                                 20021126
                                             JP 2000-606264
                                                                    20000317 <--
    NZ 513840
                                20040227
                                             NZ 2000-513840
                                                                    20000317 <--
                          Α
                                             NZ 2001-513842
                                                                    20010317 <--
     NZ 513842
                          Α
                                20040528
    NO 2001004325
                          Α
                                20011114
                                             NO 2001-4325
                                                                    20010905 <--
     ZA 2001007638
                                20020611
                                             ZA 2001-7638
                                                                    20010917 <--
                          Α
     ZA 2001007637
                          Α
                                20020621
                                             ZA 2001-7637
                                                                    20010917 <--
                                             ZA 2001-7640
     ZA 2001007640
                                20020911
                                                                    20010917 <--
                          Α
PRAI GB 1999-6437
                          Α
                                19990319
     GB 1999-9077
                          Α
                                19990420
                                           <--
     GB 1999-9466
                          Α
                                19990423
     GB 1999-16677
                                 19990715
                          Α
                                20000317
     WO 2000-EP2468
CLASS
                 CLASS PATENT FAMILY CLASSIFICATION CODES
PATENT NO.
                 ICM
WO 2000056360
                        A61K039-385
                        A61K039-39; A61K039-02; A61K039-005; A61K039-116;
                 ICS
                        A61P031-04
AB
     The present invention relates to the field of bacterial polysaccharide
     antigen vaccines. In particular, the present invention relates to
     bacterial polysaccharides conjugated to protein D from H. influenzae.
ST
    bacteria polysaccharide antigen vaccine protein D
IT
    Neisseria meningitidis
        (C; bacterial polysaccharide antigen vaccine)
     Proteins, specific or class
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (CbpA or choline-binding protein A; bacterial polysaccharide antigen
        vaccine)
IT
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (D; bacterial polysaccharide antigen vaccine)
IT
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (PsaA; bacterial polysaccharide antigen vaccine)
IT
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (PspA (pneumococcal surface protein A); bacterial polysaccharide
        antigen vaccine)
```

```
IT
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (PspC; bacterial polysaccharide antigen vaccine)
IT
     Immunity
        (Th1 adjuvant; bacterial polysaccharide antigen vaccine)
IT
     Polysaccharides, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (Vi; bacterial polysaccharide antigen vaccine)
IT
     Neisseria meningitidis
        (Y; bacterial polysaccharide antigen vaccine)
IT
     Immunostimulants
        (adjuvants, Th1; bacterial polysaccharide antigen vaccine)
IT
     Haemophilus influenzae
       Immunostimulants
     Pathogen
     Salmonella typhi
     Susceptibility (genetic)
     Trypanosoma cruzi
       Vaccines
        (bacterial polysaccharide antigen vaccine)
TT
     Lipopolysaccharides
     Saponins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (bacterial polysaccharide antigen vaccine)
TT
     Infection
         (bacterial; bacterial polysaccharide antigen vaccine)
     Polysaccharides, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (capsular; bacterial polysaccharide antigen vaccine)
TΤ
     Antigens
     Polysaccharides, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (conjugates; bacterial polysaccharide antigen vaccine)
     Glycolipoproteins
IT
     Glycolipoproteins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (glycan-containing, phospho-; bacterial polysaccharide antigen vaccine)
TΤ
     Neisseria meningitidis
        (group B polysaccharide; bacterial polysaccharide antigen vaccine)
IT
     Oligosaccharides, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (lipopeptidophospho; bacterial polysaccharide antigen vaccine)
TT
     Moraxella catarrhalis
     Shigella sonnei
        (lipopolysaccharide; bacterial polysaccharide antigen vaccine)
     Mucopolysaccharides, biological studies
Mucopolysaccharides, biological studies
TT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (lipoproteoglycans, phospho-; bacterial polysaccharide antigen vaccine)
     Proteins, specific or class
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (outer surface proteins; bacterial polysaccharide antigen vaccine)
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
       (phosphoglycan; bacterial polysaccharide antigen vaccine)
IT
    Hemolysins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pneumolysins; bacterial polysaccharide antigen vaccine)
     Bacteria (Eubacteria)
IT
     Cryptococcus neoformans
     Mycobacterium
     Neisseria meningitidis
     Staphylococcus aureus
     Streptococcus agalactiae
     Streptococcus pneumoniae
        (polysaccharide; bacterial polysaccharide antigen vaccine)
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (secreted; bacterial polysaccharide antigen vaccine)
IT
     Oligonucleotides
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (stimulatory CpG-containing; bacterial polysaccharide antigen vaccine)
IT
     Toxoids
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (tetanus; bacterial polysaccharide antigen vaccine)
     7784-30-7, Aluminum phosphate 9001-50-7, Glyceraldehyde-3-phosphate
```

```
dehydrogenase 128478-31-9
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (bacterial polysaccharide antigen vaccine)
     ANSWER 5 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
L44
AN
     2000:68366 HCAPLUS
     132:127726
DN
ED
     Entered STN:
                   28 Jan 2000
ΤI
     Adjuvant and vaccine compositions containing monophosphoryl lipid A
IN
     Laposta, Vincent James; Eldridge, John Hayward
     American Cyanamid Company, USA
PA
     PCT Int. Appl., 35 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
LA
     ICM A61K039-00
TC
CC
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 15
FAN.CNT 1
                                             APPLICATION NO.
     PATENT NO.
                         KIND
                                DATE
                                                                    DATE
                                 _ _ _'_ - - - -
     WO 2000003744
                          A2
                                20000127
                                             WO 1999-US15942
                                                                    19990713 <-
PΙ
     WO 2000003744
                          АЗ
                                20000427
         W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY,
                                                      CA,
                                                          CH CN.
                                                                  CU. CZ. DE.
             DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IN,
                                                                  IS, JP,
             KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW,
             MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR,
             TT, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
             ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     CA 2333578
                                20000127
                                             CA 1999-2333578
                                                                    19990713 <--
                          AA
                                20000207
                                             AU 1999-51018
                                                                    19990713 <--
     AU 9951018
                          A1
     AU 763770
                          B2
                                20030731
     BR 9912067
                          Α
                                 20010410
                                             BR 1999-12067
                                                                    19990713 <--
     EP 1096954
                          A2
                                20010509
                                             EP 1999-935566
                                                                    19990713 <--
                                20041006
     EP 1096954
                          B1
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
                                             US 1999-352526
     US 6306404
                          В1
                                20011023
                                                                    19990713 <--
     JP 2002520373
                                20020709
                                             JP 2000-559878
                          T2
                                                                    19990713 <--
     US 2002025330
                          A1
                                20020228
                                             US 2001-943028
                                                                    20010830 <--
     US 6635261
                          В2
                                20031021
PRAI US 1998-115392
                          Α
                                19980714
     US 1998-155270P
                          Ρ
                                19980714
                                           <--
     US 1999-352526
                                           <--
                          A3
                                19990713
     WO 1999-US15942
                          W
                                19990713
CLASS
PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
                 ----
WO 2000003744
                ICM
                        A61K039-00
                 ECLA
                        A61K039/39
US 2002025330
AΒ
    The invention pertains to adjuvant and vaccine compns. of monophosphoryl
     lipid A, sugar, and optionally an amine-based surfactant, which when
     frozen and thawed or lyophilized and reconstituted reform a colloidal
     suspension having a light transmission of greater than or equal to 88 % as
     measured spectrophotometrically.
ST
    vaccine adjuvant formulation monophosphoryl lipid A
IT
     Chlamydia
     Colloids
      Haemophilus influenzae
     Helicobacter pylori
    Human herpesvirus
     Human immunodeficiency virus
     Human papillomavirus
     Human parainfluenza virus
     Influenza virus
     Measles virus
      Moraxella catarrhalis
     Neisseria gonorrhoeae
    Neisseria meningitidis
     Norwalk virus
     Optical absorption
     Respiratory syncytial virus
     Rotavirus
     Salmonella typhi
```

```
Solvents
     Spectrophotometry
     Streptococcus group A
     Streptococcus group B
     Streptococcus pneumoniae
     Turbidimetry
        Vaccines
         (adjuvant and vaccine compns. containing monophosphoryl lipid A)
     Allergens
IT
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU
      (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (adjuvant and vaccine compns. containing monophosphoryl lipid A)
     Carbohydrates, biological studies
IT
     RL: PEP (Physical, engineering or chemical process); THU (Therapeutic
     use); BIOL (Biological study); PROC (Process); USES (Uses)
         (adjuvant and vaccine compns. containing monophosphoryl lipid A)
     Immunostimulants
         (adjuvants; adjuvant and vaccine compns. containing
         monophosphoryl lipid A)
IT
     Surfactants
         (amine-based; adjuvant and vaccine compns. containing monophosphoryl lipid
         A)
     Bacteria (Eubacteria)
TΤ
     Neoplasm
     Parasite
     Virus
         (antigens of; adjuvant and vaccine compns. containing monophosphoryl lipid
         A)
     Polysaccharides, biological studies
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (capsular; adjuvant and vaccine compns. containing monophosphoryl lipid A)
     Drug delivery systems
         (carriers; adjuvant and vaccine compns. containing monophosphoryl lipid A)
TT
     Physiological saline solutions
         (diluent; adjuvant and vaccine compns. containing monophosphoryl lipid A)
IT
     Drug delivery systems
         (freeze-dried; adjuvant and vaccine compns. containing monophosphoryl lipid
         A)
IT
     Lipopolysaccharides
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU
      (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (of Salmonella minnesota R595; adjuvant and vaccine compns. containing
         monophosphoryl lipid A)
     7784-30-7, Aluminum phosphate
                                         220048-47-5
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (adjuvant and vaccine compns. containing monophosphoryl lipid A)
     50-99-7, Dextrose, biological studies 57-48-7, D-Fructose, biological studies 57-50-1, biological studies 59-23-4, Galactose, biological
     studies 57-50-1, biological studies 59-23-4, Galactose, biologistudies 63-42-3, Lactose 69-79-4, Maltose 99-20-7, Trehalose
     499-40-1, Isomaltose
                               3458-28-4, Mannose
     RL: PEP (Physical, engineering or chemical process); THU (Therapeutic
     use); BIOL (Biological study); PROC (Process); USES (Uses)
         (adjuvant and vaccine compns. containing monophosphoryl lipid A)
     7732-18-5, Water, uses
     RL: NUU (Other use, unclassified); USES (Uses)
         (diluent; adjuvant and vaccine compns. containing monophosphoryl lipid A)
     102-71-6, biological studies 121-44-8, biological studies RL: MOA (Modifier or additive use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
         (surfactant; adjuvant and vaccine compns. containing monophosphoryl lipid
L44 ANSWER 6 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
AN
     1999:626070 HCAPLUS
     131:262583
DN
ED
     Entered STN: 01 Oct 1999
ΤI
     Haemophilus influenzae B-DTPa combination vaccine
     Artois, Claude; De Heyder, Koen; Desmons, Pierre; Garcon, Nathalie;
```

```
Mainil, Roland
     Smithkline Beecham Biologicals SA, Belg.
PA
     PCT Int. Appl., 36 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
     ICM A61K039-295
IC
         A61K039-39; A61K039-05; A61K039-08; A61K039-09; A61K039-095;
     ICS
          A61K039-10; A61K039-102; A61K039-13; A61K039-29
CC
     63-3 (Pharmaceuticals)
FAN.CNT 1
                                DATE
                         KIND
                                            APPLICATION NO.
                                                                    DATE
     PATENT NO.
                         ----
     WO 9948525
                          Al
                                19990930
                                            WO 1999-EP1959
                                                                    19990322 <--
         W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS,
             JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,
             MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
             TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
             MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
             ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                            CA 1999-2325436
     CA 2325436
                          AA
                                19990930
                                                                    19990322 <--
                                19991018
                                            AU 1999-34172
                                                                    19990322 <--
     AU 9934172
                          Α1.
     AU 735619
                          B2
                                20010712
                                20001205
                                             BR 1999-9037
                                                                    19990322 <--
     BR 9909037
                          Α
     TR 200002737
                          T2
                                20001221
                                            TR 2000-200002737
                                                                    19990322 <--
                                            EP 1999-915692
                                                                    19990322 <--
                          A1
                                20010110
     EP 1066053
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, FI
     JP 2002507581
                          T2
                                20020312
                                             JP 2000-537572
                                                                    19990322 <--
                                            NZ 1999-506604
                                20030228
                                                                    19990322 <--
     NZ 506604
                          Α
     ZA 2000004956
                          Α
                                20020108
                                            ZA 2000-4956
                                                                    20000918 <--
     NO 2000004758
                                20001108
                                            NO 2000-4758
                                                                    20000922 <--
     US 2003022304
                          A1
                                20030130
                                            US 2002-217572
                                                                    20020813 <--
PRAT GR 1998-6456
                                19980325
                          А
     WO 1999-EP1959
                          W
                                19990322
                                          <--
     US 2000-647032
                          В1
                                20001031
CLASS
                 CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
 -----
                 ----
                        _____
 WO 9948525
                 ICM
                        A61K039-295
                 ICS
                        A61K039-39; A61K039-05; A61K039-08; A61K039-09;
                        A61K039-095; A61K039-10; A61K039-102; A61K039-13;
                        A61K039-29
 US 2003022304
                 ECLA
                        A61K039/05; A61K039/08; A61K039/09; A61K039/095;
                        A61K039/10; A61K039/102; A61K039/13; A61K039/29;
                        A61K039/39
    This invention relates to a general method by which either
AΒ
     extemporaneously prepared or liquid Haemophilus influenzae B (Hib)/DTPa
     combination vaccines can be made in order to avoid Hib interference while
     being able to maintain the maximum, stable adsorption of each antigen onto
     the aluminum-based adjuvant on which it is most immunogenic. In so doing,
     pertussis antigens in combination vaccines of the present invention are
     stably retained in their most potent form. Examples are given for the
     vaccines using Al hydroxide or Al phosphate as adjuvants.
ST
     vaccine Haemophilus diphtheria tetanus pertussis
TT
     Vaccines
        (Haemophilus influenzae B-DTPa combination vaccine)
     Antigens
     Polysaccharides, biological studies
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); THU
     (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses)
        (Haemophilus influenzae B-DTPa combination vaccine)
     Immunostimulants
        (adjuvants; Haemophilus influenzae B-DTPa combination
        vaccine)
IT
    Hepatitis A virus
     Human poliovirus
        (antigens; Haemophilus influenzae B-DTPa combination vaccine)
TT
     Streptococcus pneumoniae
        (capsular polysaccharide and proteins; Haemophilus influenzae B-DTPa
        combination vaccine)
TT
     Toxoids
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
```

Search done by Noble Jarrell

Graser 10/030313 Page 18

```
(diphtheria; Haemophilus influenzae B-DTPa combination vaccine)
IT
    Neisseria meningitidis
        (group A, capsular polysaccharide; Haemophilus influenzae B-DTPa
        combination vaccine)
     Neisseria meningitidis
IT
        (group C, capsular polysaccharide; Haemophilus influenzae B-DTPa
        combination vaccine)
IT
     Antigens
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hepatitis B surface; Haemophilus influenzae B-DTPa combination
        vaccine)
IT
    Moraxella catarrhalis
        (outer membrane proteins; Haemophilus influenzae B-DTPa combination
IT
     Toxoids
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pertussis; Haemophilus influenzae B-DTPa combination vaccine)
     Toxoids
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (tetanus; Haemophilus influenzae B-DTPa combination vaccine)
IT
     Haemophilus influenzae
        (type b; Haemophilus influenzae B-DTPa combination vaccine)
                                   21645-51-2, Aluminum hydroxide, biological
IT
     7784-30-7, Aluminum phosphate
     studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (Haemophilus influenzae B-DTPa combination vaccine)
             THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 6
RE
(1) Corbel, M; BIOLOGICALS 1994, V22(4), P353 HCAPLUS
(2) Corbel, M; BIOLOGICALS 1997, V25/3, P351
(3) Ellis, R; VACCINE 1999, V17(13-14), P1635 MEDLINE
(4) Slaoui Moncef Mohamed; WO 9746255 A 1997 HCAPLUS
(5) Smithkline Beecham Biolog; WO 9324148 A 1993 HCAPLUS
(6) Smithkline Beecham Biolog; WO 9700697 A 1997 HCAPLUS
L44 ANSWER 7 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
    1999:597423 HCAPLUS
AN
DN
    131:213104
                  22 Sep 1999
ED
     Entered STN:
    Antiquenic conjugates of conserved lipopolysaccharides of gram negative
ΤI
    bacteria
IN
    Arumugham, Rasappa G.; Fortuna-Nevin, Maria; Apicella, Michael A.; Gibson,
    Bradford W.
PA
    American Cyanamid Company, USA
    Eur. Pat. Appl., 18 pp.
SO
    CODEN: EPXXDW
DT
    Patent
LA
    English
IC
    ICM A61K039-385
ICI A61K039-02, A61K039-095
    15-2 (Immunochemistry)
     Section cross-reference(s): 14, 63
FAN.CNT 1
    PATENT NO.
                               DATE
                                           APPLICATION NO.
                                                                  DATE
                        KIND
                               -----
     -----
                                           -----
                                                                  -----
                        _ _ _ _
    EP 941738
                         A1
                               19990915
                                          EP 1999-301747
                                                                  19990309 <--
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO
                               19990910
    CA 2264970
                                           CA 1999-2264970
                                                                  19990308 <--
                         AΑ
    AU 9919540
                         A1
                               19990923
                                           AU 1999-19540
                                                                  19990309 <--
    AU 766184
                         B2
                               20031009
    JP 11322793
                         A2
                               19991124
                                           JP 1999-61354
                                                                  19990309 <--
    BR 9902008
                               20000509
                                          BR 1999-2008
                                                                  19990309 <--
                         Α
PRAI US 1998-37529
                         Α
                               19980310 <--
CLASS
PATENT NO.
                CLASS PATENT FAMILY CLASSIFICATION CODES
EP 941738
                TCM
                       A61K039-385
                ICI
                       A61K039-02, A61K039-095
    Antigenic conjugates are provided which comprise a carrier protein
    covalently bonded to the conserved portion of a lipopolysaccharide of a
    gram neg. bacteria, wherein said conserved portion of the
    lipopolysaccharide comprises the inner core and lipid A portions of said
     lipopolysaccharide, said conjugate eliciting a cross reactive immune
    response against heterologous strains of said gram neg. bacteria. The
```

carrier protein is selected from CRM197, tetanus toxin, diphtheria toxin,

pseudomonas exotoxin A, cholera toxin, group A streptococcal toxin, pneumolysin of Streptococcus pneumoniae, filamentous hemagglutinin (FHA), FHA of Bordetella pertussis, pili or pilins of Neisseria gonorrhoeae or meningitidis, outer membrane proteins of Neisseria meningitidis, C5A peptidase of Streptococcus and surface protein of Moraxella catarrhalis. gram neg bacteria lipopolysaccharide carrier protein; vaccine lipopolysaccharide carrier conjugate immune adjuvant TT Hemagglutinins RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (FHA (filamentous hemagglutinin); conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune response against heterologous strains of gram neg. bacteria) IT Proteins, specific or class RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (OMP (outer membrane protein), carrier; conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune response against heterologous strains of gram neg. bacteria) Proteins, specific or class IT RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (SU (surface), carrier; conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune response against heterologous strains of gram neg. bacteria) IT Immunostimulants (adjuvants; conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune response against heterologous strains of gram neg. bacteria) TТ Toxins RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (bacterial; conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune response against heterologous strains of gram neg. bacteria) IT Streptococcus (carrier; conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune response against heterologous strains of gram neg. bacteria) TΤ Toxins RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (cholera; conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune response against heterologous strains of gram neg. bacteria) IT Bordetella Bordetella pertussis Chlamydia Escherichia coli Gram-negative bacteria Haemophilus Haemophilus ducreyi Haemophilus influenzae Helicobacter pylori Klebsiella Moraxella catarrhalis Neisseria Neisseria gonorrhoeae Neisseria meningitidis Pilus Proteus mirabilis Pseudomonas Pseudomonas aeruginosa Salmonella Salmonella minnesota Salmonella typhimurium Shigella Streptococcus pneumoniae Vaccines Vibrio cholerae (conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune response against heterologous strains of gram neg. bacteria) IT Lipid A

```
Pilins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (conjugates of conserved lipopolysaccharides of gram neg. bacteria and
        carrier proteins for eliciting cross reactive immune response against
        heterologous strains of gram neg. bacteria)
     Lipopolysaccharides
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (conjugates of conserved lipopolysaccharides of gram neg. bacteria and
        carrier proteins for eliciting cross reactive immune response against
        heterologous strains of gram neg. bacteria)
IT
     Carriers
        (conjugates; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (conjugates; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
     Toxins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (diphtheria, carrier; conjugates of conserved lipopolysaccharides of
        gram neg. bacteria and carrier proteins for eliciting cross reactive
        immune response against heterologous strains of gram neg. bacteria)
TT
     Toxins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (exotoxin A, carrier; conjugates of conserved lipopolysaccharides of
        gram neg. bacteria and carrier proteins for eliciting cross reactive
        immune response against heterologous strains of gram neg. bacteria)
IT
     Drug delivery systems
        (injections, i.m.; conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
     Drug delivery systems
        (injections, i.v.; conjugates of conserved lipopolysaccharides of gram
        neg. bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
        (injections, s.c.; conjugates of conserved lipopolysaccharides of gram neg. bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
     Drug delivery systems
        (intradermal; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
     Drug delivery systems
        (nasal, intra-; conjugates of conserved lipopolysaccharides of gram
        neg. bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
     Drug delivery systems
        (ophthalmic; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
TT
     Drug delivery systems
        (oral; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
     Hemolysins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (pneumolysins, carrier; conjugates of conserved lipopolysaccharides of
        gram neg. bacteria and carrier proteins for eliciting cross reactive
        immune response against heterologous strains of gram neg. bacteria)
TT
     Drug delivery systems
        (solns., i.p.; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
     Toxins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (tetanus, carrier; conjugates of conserved lipopolysaccharides of gram
```

O antigen

Graser 10/030313 Page 21

```
neg. bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
     Streptococcus group A
        (toxin; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
IT
     Drug delivery systems
        (vaginal; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
     100179-39-3, C5A Peptidase
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (carrier; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
     response against heterologous strains of gram neg. bacteria)
51-85-4, Cystamine 1071-93-8, Adipic acid dihydrazide 1892-57-5, EDAC
TT
     6539-14-6, Traut's reagent 42014-51-7, Bromoacetic acid
N-hydroxysuccinimide ester 57757-57-0 64202-52-4 64987-85-5, SMCC
     68181-17-9, SPDP 72252-96-1, SIAB 76931-93-6, SATA 79886-55-8,
     Succinimidyl 4-(p-maleimidophenyl)butyrate 150205-95-1 150244-18-1
     158913-22-5
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (linker; conjugates of conserved lipopolysaccharides of gram neg.
        bacteria and carrier proteins for eliciting cross reactive immune
        response against heterologous strains of gram neg. bacteria)
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 3
RE
(1) Baker, P; Infection and Immunity 1994, V62(6), P2257 HCAPLUS (2) Rune, A; Microbial Pathogenesis 1997, V23(3), P139
(3) Stanislavsky, E; FEMS Microbiology Reviews 1997, V21(3), P243 HCAPLUS
L44 ANSWER 8 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
     1999:571730 HCAPLUS
AN
     131:213099
     Entered STN: 09 Sep 1999
ED
     Vaccine for Moraxella catarrhalis
TT
IN
     Murphy, Timothy F.
     The Research Foundation of State University of New York, USA
PA
     U.S., 20 pp., Cont.-in-part of U.S. 5,607,846.
so
     CODEN: USXXAM
DT
     Patent
LΑ
     English
     ICM A61K039-02
ICS C07K014-00
IC
NCL 424251100
     15-2 (Immunochemistry)
     Section cross-reference(s): 3
FAN.CNT 2
                          KIND DATE
                                                                       DATE
     PATENT NO.
                                              APPLICATION NO.
                          ----
     -----
                                  -----
                                               -----
                                                                       ------
     US 5948412
                                  19990907
                                              US 1997-810655
                                                                       19970303 <--
                           Α
     US 5607846
                          Α
                                  19970304
                                              US 1994-245758
                                                                       19940517 <--
                           AA
                                  19951123
                                              CA 1995-2189971
                                                                       19950420 <--
     CA 2189971
                           С
     CA 2189971
                                  20030729
     ES 2202361
                           Т3
                                  20040401
                                              ES 1995-917165
                                                                      19950420 <--
PRAI US 1994-245758
                                 19940517 <--
CLASS
                  CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
 US 5948412
                 ICM
                         A61K039-02
                         C07K014-00
                 ICS
                         424251100
                 NCL
 US 5948412
                 ECLA
                        C07K014/21B
                                                                                 <-- .
 US 5607846
                 ECLA
                         C07K014/21B
    Compns. comprising outer membrane protein E, and peptides and
     oligopeptides thereof, of Moraxella catarrhalis are described. Addnl.,
     nucleotide sequences encoding the protein, peptide, or oligopeptide are
     disclosed, as well as recombinant vectors containing these sequences.
     Protein, peptide, or oligopeptide can be produced from host cell systems
     containing these recombinant vectors. Peptides and oligopeptides can also be chemical synthesized. Disclosed are the uses of the protein, peptides and
     oligopeptides as antigens in antigenic formulations for vaccine
     applications or for generating antisera of diagnostic or therapeutic use;
```

and as antigens in diagnostic immunoassays. The nucleotide sequences are useful for constructing vectors for use as vaccines for insertions into

Graser 10/030313 attenuated bacteria in constructing a recombinant bacterial vaccine and for inserting into a viral vector in constructing a recombinant viral vaccine. Also described is the use of nucleotide sequences related to the gene encoding E as primers and/or probes in mol. diagnostic assays for the detection of M. catarrhalis. Moraxella catarrhalis outer membrane protein E; vaccine antiserum outer membrane protein E; gene protein E epitope probe primer Primers (nucleic acid) TT Primers (nucleic acid) RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (DNA; vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis) Proteins, specific or class RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (OMP (outer membrane protein), E; vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis) Polysaccharides, biological studies TТ RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (capsular; vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis) Drug delivery systems (carriers; vaccine for Moraxella catarrhalis comprising outer membrane IT protein E epitope and primers and probes for genetic diagnosis) IT Diagnosis (immunodiagnosis; vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis) DNA IT DNA RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (primer; vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis) Animal cell line Antiserums Bacteria (Eubacteria) DNA sequences Epitopes Filamentous fungi Haemophilus influenzae Insect (Insecta)
Mammal (Mammalia) Molecular cloning

Page 22

Moraxella catarrhalis Neisseria meningitidis Protein sequences Pseudomonas aeruginosa Staphylococcus aureus Streptococcus pneumoniae Vaccines

Virus vectors Yeast

(vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis)

Gene, microbial RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis)

Antigens

Lipopolysaccharides

Polysaccharides, biological studies

RNA

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis)

Probes (nucleic acid)

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

Graser 10/030313

Page 23

(vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis) 159869-80-4 174065-50-0 242799-79-7 TТ RL: PRP (Properties) (amino acid sequence; vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis) TT 174066-42-3 RL: PRP (Properties) (nucleotide sequence; vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis) IT 173432-99-0 173433-06-2 173433-09-5 173433-10-8 173433-11-9 173433-12-0 173433-13-1 RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (vaccine for Moraxella catarrhalis comprising outer membrane protein E epitope and primers and probes for genetic diagnosis) THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT RE (1) Bartos; J Infect Dis 1988, V158, P761 HCAPLUS (2) Bhushan; Abstracts Gen Meet Am Soc Microbiol 1991, V97, P30 (3) Maciver; J Infect Dis 1993, V168, P469 MEDLINE L44 ANSWER 9 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN AN 1999:487519 HCAPLUS 131:120851 ED Entered STN: 06 Aug 1999 Nonrecombinant subunit vaccine TI Gerlach, Gerald-F.; Goethe, Ralph IN PA Germany Ger. Offen., 22 pp. SO CODEN: GWXXBX DTPatent LΑ German IC ICM A61K039-102 ICS A61K039-095 CC 63-4 (Pharmaceuticals) Section cross-reference(s): 15 FAN.CNT 1 PATENT NO. APPLICATION NO. KIND DATE DATE -------------------------DE 19753176 A1 19990729 DE 1997-19753176 19971120 <--DE 19753176 C2 20000427 PRAI DE 1997-19753176 19971120 <--CLASS PATENT NO. CLASS · PATENT FAMILY CLASSIFICATION CODES DE 19753176 ICM A61K039-102 A61K039-095 TCS The title bacterial vaccines are obtained by (1) cultivation of (preferably gram-neg.) pathogenic bacteria, preferably under mineral or nutrient deficiency stress or heat stress, and (2) enrichment of protective antigens from the bacteria by use of detergents, especially steroidal detergents such as cholic acid. This procedure exts. various protective antigens (especially lipoproteins) from the outer membrane without lysing the bacteria and thus without causing release of extraneous proteins. The subunit vaccine can be used as a marker vaccine for differentiation of vaccinated from infected subjects by ELISA. Thus, Actinobacillus pleuropneumoniae 811/051 (serotype 9) was cultivated in PPLO medium + Iso Vitale X at 37.degree. under Fe deficiency conditions (100 .mu.M 2,2'-dipyridyl), centrifuged, and resuspended in distilled water, and transferrin-binding protein A was extracted from the outer membrane with 0.075% Na deoxycholate. This extract and a similar extract from serotype 2 were combined 1:2, diluted 1:10, and mixed with HCHO 0.05 and Emulsigen Plus 20% for use as a vaccine in swine. ST bacteria vaccine outer membrane protein; Actinobacillus vaccine detergent extn; pleuropneumonia vaccine extn deoxycholate Proteins, specific or class RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (OMP (outer membrane protein); nonrecombinant subunit vaccine) IT Detergents (anionic; nonrecombinant subunit vaccine) Mineral elements, biological studies

```
RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (bacteria deficiency in; nonrecombinant subunit vaccine)
     Nutrition, microbial
IT
         (deficiency; nonrecombinant subunit vaccine)
TΤ
     Immunoassav
        (enzyme-linked immunosorbent assay, for bacterial outer membrane
        proteins; nonrecombinant subunit vaccine)
IT
        (immunization of, against pleuropneumonia; nonrecombinant subunit
        vaccine)
IT
     Diagnosis
         (immunodiagnosis, of pleuropneumonia; nonrecombinant subunit vaccine)
     Detergents
IT
         (nonionic; nonrecombinant subunit vaccine)
IT
     Actinobacillus equuli
     Actinobacillus pleuropneumoniae
     Chelating agents
     Detergents
     Escherichia coli
     Gram-negative bacteria
       Haemophilus actinomycetemcomitans
     Haemophilus agni
       Haemophilus influenzae
       Haemophilus paragallinarum
       Haemophilus parasuis
     Haemophilus somnus
     Mannheimia haemolytica
       Moraxella bovis
       Moraxella catarrhalis
       Moraxella lacunata
     Neisseria gonorrhoeae
     Neisseria meningitidis
     Neisseriaceae
     Pasteurella avium
     Pasteurella multocida
     Pasteurellaceae
     Stress, microbial
       Vaccines
        (nonrecombinant subunit vaccine)
     Antigens
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PUR (Purification or recovery); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (nonrecombinant subunit vaccine)
     Bile acids
     Bile salts
     Quaternary ammonium compounds, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (nonrecombinant subunit vaccine)
     Antibodies
     RL: ANT (Analyte); BAC (Biological activity or effector, except adverse);
     BSU (Biological study, unclassified); THU (Therapeutic use); ANST
     (Analytical study); BIOL (Biological study); USES (Uses)
        (to Actinobacillus pleuropneumoniae; nonrecombinant subunit vaccine)
     Proteins, specific or class
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PUR (Purification or recovery); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (transferrin-binding; nonrecombinant subunit vaccine)
IT
     Nutrition, animal
        (undernutrition; nonrecombinant subunit vaccine)
TT
     Detergents
        (zwitterionic; nonrecombinant subunit vaccine)
IT
     7439-89-6, Iron, biological studies
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
     (chelating agents for; nonrecombinant subunit vaccine) 60-00-4, EDTA, biological studies 67-43-6D, salts 70-
IT
                                                              70-51-9.
     Deferrioxamine 139-13-9 366-18-7, 2,2'-Dipyridyl 12111-24-9
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (nonrecombinant subunit vaccine)
                                                 81-24-3, Taurocholic acid
     57-09-0, Cetyltrimethylammonium bromide
     81-25-4, Cholic acid 83-44-3, Deoxycholic acid 98-11-3D,
Benzenesulfonic acid, alkyl derivs., biological studies 151-21-3, SDS,
     biological studies 302-95-4, Sodium deoxycholate
                                                            360-65-6,
```

Graser 10/030313

Page 25

```
Glycodeoxycholic acid 361-09-1, Sodium cholate 475-31-0, Glycocholic acid 516-50-7, Taurodeoxycholic acid 2044-56-6, Lithium dodecyl
     sulfate 7631-98-3, Sodium N-laurylsarcosinate 9002-92-0, Brij 35
     9002-93-1, Triton X-100 9004-95-9, Brij 58 9005-64-5, Tween 20 9005-65-6, Tween 80 9036-19-5, Nonidet P40 9043-30-5, Genapol X-080
     14933-09-6, N-Tetradecyl-N,N-dimethyl-3-ammonio-1-propanesulfonate
     25339-99-5, Sucrose monolaurate 29836-26-8 59122-55-3 68894-53-1,
     Tergitol 69227-93-6 75621-03-3, CHAPS 82473-24-3, CHAPSO 85261-20-7, MEGA 10 85316-98-9, MEGA 8 86295-19-4,
     N-Dodecyl-N, N-dimethyl-3-ammonio-1-propanesulfonate 86303-23-3, Deoxy
     BigCHAP 106392-12-5, Synperonic PE/F 68 232601-34-2, Tween 48
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (nonrecombinant subunit vaccine)
RE.CNT
              THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
(1) Anon; DD 242716 A3 HCAPLUS
(2) Anon; EP 37931 A2 HCAPLUS
(3) Anon; US 4845036 HCAPLUS
(4) Wpids; AU 9061356 A HCAPLUS
(5) Wpids; AU 9534351 A
L44 ANSWER 10 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
     1998:800024 HCAPLUS
AN
DN
     130:51336
     Entered STN: 22 Dec 1998
ED
TI
     Laft mutants of pathogenic gram-negative bacteria
     Apicella, Michael A.; Gibson, Bradford W.; Nichols, Wade A.
PA
     University of Iowa Research Foundation, USA; University of California
     PCT Int. Appl., 31 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
     ICM A61K039-02
ICS A01N063-00; C12N001-00; C12N001-20
IC
CC
     15-2 (Immunochemistry)
     Section cross-reference(s): 3, 10
FAN.CNT 1
                                                                        DATE
     PATENT NO.
                          KIND DATE
                                               APPLICATION NO.
     _____
                           ____
                                  _____
                                                ______
                                                                        _____
     WO 9853851
                                  19981203 WO 1998-US10881
                           A1
                                                                       19980528 <--
ΡI
         W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG,
              KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
              UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
              CM, GA, GN, ML, MR, NE, SN, TD, TG
                           A1 19981230 AU 1998-77010
     AU 9877010
                                                                        19980528 <--
PRAI US 1997-47791P
                           P
                                  19970528 <--
     WO 1998-US10881
                           W
                                  19980528 <--
CLASS
 PATENT NO.
                  CLASS PATENT FAMILY CLASSIFICATION CODES
                 ICM A61K039-02
 WO 9853851
                         A01N063-00; C12N001-00; C12N001-20
                  ICS
 WO 9853851
                  ECLA
                         C12N009/10C1
AB A method is provided for identifying, isolating, and producing
     lipooligosaccharide (LOS) mutants of gram-neg. bacterial pathogens. The
     method comprises mutating the laft gene of a gram-neg. bacterial pathogen
     so that there is a lack of a functional Lipid A fatty acid transferase
     protein. The resulting LOS mutants lack one or more secondary acyl chains
     as compared to the LOS contained in the wild type gram-neg. bacterial
     pathogen. The LOS isolated from the laft mutants displays substantially
     reduced toxicity as compared to that of the wild type strain. Also, the
     present invention provides methods for using a vaccine formulation containing
     the laft mutants, the endotoxin isolated therefrom, or the endotoxin
     isolated therefrom which is then conjugated to a carrier protein, to
     immunize an individual against infections caused by gram-neg. bacterial
     pathogens by administering a prophylactically effective amount of the
     vaccine formulation.
ST
    lipid A fatty acid transferase gene; lipopolysaccharide endotoxin vaccine
     gram neg bacteria
TT
     Immunostimulants
        (adjuvants; bacterium having mutated lipid A fatty acid
        transferase gene as vaccine for preventing infections by pathogenic
```

```
gram-neg. bacteria)
IT
     Microorganism
        (antiqen; bacterium having mutated lipid A fatty acid transferase gene
        as vaccine for preventing infections by pathogenic gram-neg. bacteria)
IT
     Infection
        (bacterium having mutated lipid A fatty acid transferase gene as
        vaccine for preventing infections by pathogenic gram-neg. bacteria)
     Drug delivery systems
IT
        (carriers; bacterium having mutated lipid A fatty acid transferase gene
        as vaccine for preventing infections by pathogenic gram-neg. bacteria)
IT
     Toxins
     RL: PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (endotoxins; gram-neg. bacterium having mutated lipid A fatty acid
        transferase gene as vaccine for preventing infections)
     Bordetella pertussis
     Escherichia coli
     Gram-negative bacteria
       Haemophilus ducreyi
       Haemophilus influenzae
     Human adenovirus
     Human parainfluenza virus
     Influenza virus
     Legionella pneumophila
       Moraxella catarrhalis
     Mycoplasma pneumoniae
     Neisseria gonorrhoeae
     Pneumocystis carinii
     Pseudomonas aeruginosa
     Respiratory syncytial virus
     Streptococcus group A
     Streptococcus pneumoniae
       Vaccines
        (gram-neg. bacterium having mutated lipid A fatty acid transferase gene
        as vaccine for preventing infections)
     Antigens
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (gram-neg. bacterium having mutated lipid A fatty acid transferase gene
        as vaccine for preventing infections)
     Lipopolysaccharides
     RL: PUR (Purification or recovery); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (gram-neg. bacterium having mutated lipid A fatty acid transferase gene
        as vaccine for preventing infections)
TT
     Drug delivery systems
        (injections, i.m.; bacterium having mutated lipid A fatty acid
        transferase gene as vaccine for preventing infections by pathogenic
        gram-neg. bacteria)
     Drug delivery systems
IT
        (injections, i.v.; bacterium having mutated lipid A fatty acid
        transferase gene as vaccine for preventing infections by pathogenic
        gram-neg. bacteria)
IT
     Drug delivery systems
        (injections, s.c.; bacterium having mutated lipid A fatty acid
        transferase gene as vaccine for preventing infections by pathogenic
        gram-neg. bacteria)
     Drug delivery systems
        (intradermal; bacterium having mutated lipid A fatty acid transferase
        gene as vaccine for preventing infections by pathogenic gram-neg.
        bacteria)
     Gene, microbial
     RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
     THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
        (laft (lipid A fatty acid transferase); gram-neg. bacterium having
        mutated lipid A fatty acid transferase gene as vaccine for preventing
        infections)
TT
     Gene, microbial
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (msbB; gram-neq. bacterium having mutated lipid A fatty acid
        transferase gene as vaccine for preventing infections)
IT
     Drug delivery systems
        (mucosal; bacterium having mutated lipid A fatty acid transferase gene
        as vaccine for preventing infections by pathogenic gram-neg. bacteria)
IT
     Drug delivery systems
```

Graser 10/030313 Page 27

(nasal, intra-; bacterium having mutated lipid A fatty acid transferase gene as vaccine for preventing infections by pathogenic gram-neg. bacteria)

IT Gene

> RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(open reading frame; gram-neg. bacterium having mutated lipid A fatty acid transferase gene as vaccine for preventing infections)

IT Drug delivery systems

(ophthalmic; bacterium having mutated lipid A fatty acid transferase gene as vaccine for preventing infections by pathogenic gram-neg. bacteria)

IT Drug delivery systems

(oral; bacterium having mutated lipid A fatty acid transferase gene as vaccine for preventing infections by pathogenic gram-neg. bacteria)

IT Drug delivery systems

(solns., i.p.; bacterium having mutated lipid A fatty acid transferase gene as vaccine for preventing infections by pathogenic gram-neg. bacteria)

115926-32-4 IT

> RL: BSU (Biological study, unclassified); BIOL (Biological study) (gram-neg. bacterium having mutated lipid A fatty acid transferase gene as vaccine for preventing infections)

TΤ 217181-27-6

RL: PRP (Properties)

(nucleotide sequence; gram-neg. bacterium having mutated lipid A fatty acid transferase gene as vaccine for preventing infections)

RE.CNT THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

- (1) Clementz; J Biol Chem 1997, V272(16), P10353 HCAPLUS
- (2) Jones; Infect Immun 1997, V65(11), P4778 HCAPLUS
- (3) Lee; J Biol Chem 1995, V270(45), P27151 HCAPLUS
- (4) Somerville; J Clin Invest 1996, V97(2), P359 HCAPLUS (5) Sprouse; US 5641492 A 1997 HCAPLUS
- (6) Sunshine; J Bacteriol 1997, V179(17), P5521 HCAPLUS
- L44 ANSWER 11 OF 18 'HCAPLUS COPYRIGHT 2004 ACS on STN
- ΑN 1993:17456 HCAPLUS
- DN 118:17456
- Entered STN: 24 Jan 1993 ED
- Use of the purA gene as a selectable marker in stabilization and TΙ integration of plasmid or bacteriophage cloning vectors
- IN Brey, Robert Newton, III; Fulginiti, James Peter; Anilionis, Algis
- PA American Cyanamid Co., USA
- Eur. Pat. Appl., 29 pp. SO
- CODEN: EPXXDW
- DT Patent
- English LΑ
- ICM C12N015-74 IC
  - ICS A61K039-112
- ICI C12N015-74, C12R001-42
- 3-2 (Biochemical Genetics) Section cross-reference(s): 10, 15

FAN CNT 1

PAN. CNI I					
	PATENT NO.			APPLICATION NO.	DATE .
ΡI	EP 512260	A2	19921111	EP 1992-105887	19920406 <
	EP 512260	A3	19930728		
	R: AT, BE, CH,	DE, DK	, ES, FR,	GB, GR, IT, LI, LU, NL,	PT, SE
	AT 202800	E	20010715	AT 1992-105887	19920406 <
	ES 2160573	T3	20011116	ES 1992-105887	19920406 <
	PT 512260	T	20011228	PT 1992-105887	19920406 <
	JP 05192161	A2	19930803	JP 1992-134375	19920428 <
	JP 3320095	B2	20020903		
	NO 9201729	Α	19921104	NO 1992-1729	19920430 <
	CA 2067862	AA	19921104	CA 1992-2067862	19920501 <
	CA 2067862	C	20031230		
	AU 9215959	A1	19921105	AU 1992-15959	19920501 <
	AU 654347	B2	19941103		
	US 5919663	Α	19990706	US 1995-380297	19950130 <
	US 5961983	A	19991005	US 1995-448907	19950524 <
	GR 3036487	<b>T</b> 3	20011130	GR 2001-401346	20010831 <
PRA1	US 1991-695706	A	19910503	<	
	US 1994-204903	B1	19940302	<	
	US 1995-380297	A3	19950130	<	
CLASS					

purA gene also acts as a site for integration of the plasmid. The use of these vectors does not involve the use of antibiotic resistance markers and is therefore particularly suitable for hosts used in live vaccines. pUC8-based plasmid carrying the Escherichia coli purA gene and the gene for the nontoxic subunit of the E. coli heat-labile enterotoxin was constructed and introduced into Salmonella dublin, S. typhimurium or Salmonella vaccine strains carrying deletions in the purA gene and transformants selected on minimal medium. This plasmid was maintained in cultures grown on a minimal medium without loss for 80 generations but lost rapidly in the absence of selection (1% retention in 40 generations). When the purA gene was used in integrating vectors the prototrophic phenotype was 100% stable for at least 80 generations in the presence or absence of selection.

ST purA gene selectable marker cloning vector; Salmonella adenine auxotrophy selectable marker

IT Bordetella pertussis Chlamydia trachomatis Clostridium tetani Corynebacterium diphtheriae Escherichia coli

Haemophilus influenzae

Klebsiella pneumoniae

Moraxella catarrhalis

Neisseria gonorrhoeae

Neisseria meningitidis

Parasite

Pseudomonas aeruginosa

Staphylococcus aureus

Streptococcus pneumoniae

Streptococcus pyogenes

Vibrio cholerae

(antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in)

IT Aromatic hydrocarbons, biological studies

RL: BPN (Biosynthetic preparation); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(biosynthesis of, microorganism deficient in as host for expression vectors, complementing genes on plasmids as selectable markers for)

Plasmid and Episome

Virus, bacterial

(cloning vector, purA gene as selectable marker for, adenine auxotrophic host for)

IT Antigens

RL: BIOL (Biological study)

(genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in)

IT Campylobacter

Escherichia

Salmonella

Salmonella dublin

Salmonella typhimurium

Shigella

Vibrio

Yersinia

(in live vaccines, heterologous antigen genes in, stabilization or integration of, purA gene as selectable marker in)

Vaccines

(live, bacteria carrying cloned antigen genes for, purA gene as a selectable marker for cloning vectors in)

IT Plasmid and Episome

(pX3005, gene for heat-labile enterotoxin subunit of Escherichia coli on, integration and expression in Salmonella of, purA gene as selectable marker in)

IT Plasmid and Episome

(pX3006, gene for circumsporozoite antigen of Plasmodium berghei on, integration and expression in Salmonella of, purA gene as selectable marker in)

Plasmid and Episome IT (pX3007, chimeric gene for fusion protein of circumsporozoite antigen of Plasmodium berghei and Escherichia coli enterotoxin on, integration and expression in Salmonella of, purA gene as selectable marker in) Plasmid and Episome IT (pX3009, gene for circumsporozoite antigen of Plasmodium berghei on, integration and expression in Salmonella of, purA gene as selectable marker in) IT Plasmid and Episome (pX3010, gene for heat-labile enterotoxin subunit of Escherichia coli on, integration and expression in Salmonella of, purA gene as selectable marker in) IT Antigens RL: BIOL (Biological study) (CS (circumsporozoite), gene for, expression in enterobacteria of, in live vaccines, purA gene for stabilization or integration of antiqen genes in, fusion proteins with heat-labile enterotoxin subunit in relation to) Virus, animal IT (Epstein-Barr, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT Virus, animal (adeno-, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) ΙT Virus, animal (corona-, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) Virus, animal IT (cytomegalo-, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) RL: BIOL (Biological study) (entero-, LT, gene for, expression in enterobacteria of, in live vaccines, purA gene for stabilization or integration of enterotoxin genes in, fusion proteins with circumsporozoite antigens in relation to) TT Virus, animal (hepatitis A, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) Virus, animal IT (hepatitis B, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) Virus, animal IT (hepatitis C, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) Virus, animal IT (hepatitis, non-A, non-B, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT Virus, animal (herpes simplex 1, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT (herpes simplex 2, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT Virus, animal (human T-cell leukemia type I, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT Virus, animal (human T-cell leukemia type II, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker IT Virus, animal (human immunodeficiency 1, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT Virus, animal (human immunodeficiency 2, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT Virus, animal (influenza, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT Virus, animal (measles, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in)

IT

Virus, animal

(papilloma, antigens of, genes for, vaccine bacteria carrying, cloning

vectors using purA gene as selectable marker in)

Page 30

IT Virus, animal (parainfluenza, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT (pathogenic, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) Virus, animal IT (polio-, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) Virus, animal TT (respiratory syncytial, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) Virus, animal (rota-, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) IT Virus, animal (rubella, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) TT Virus, animal (varicella-zoster, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) TΤ Virus, animal (yellow fever, antigens of, genes for, vaccine bacteria carrying, cloning vectors using purA gene as selectable marker in) Gene, microbial RL: BIOL (Biological study) (purA, as selectable marker for cloning vectors, deletion of host copy of gene in) IT 73-24-5, Adenine, biological studies RL: BIOL (Biological study) (auxotrophy for, as selectable marker for cloning vectors, deletion of host copy of gene in) TT 120-73-0, Purine RL: BIOL (Biological study) (biosynthesis of, microorganism deficient in as host for expression vectors, complementing genes on plasmids as selectable markers for) 9023-57-8, Adenylosuccinate synthetase TT RL: BIOL (Biological study) (gene for, as selectable marker for cloning vectors, deletion of host copy of gene in) L44 ANSWER 12 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN 1991:499235 HCAPLUS 115:99235 DN Entered STN: 06 Sep 1991 ED A method for isolating and purifying transferrin and lactoferrin receptor TI proteins from bacteria and the preparation of vaccines containing the same TN Schryvers, Anthony Bernard University Technologies International Inc., Can. PA SO PCT Int. Appl., 34 pp. CODEN: PIXXD2 DTPatent English LA ICM A61K039-095 TC ICS A61K039-102; A61K039-02 CC 63-3 (Pharmaceuticals) FAN.CNT 2 PATENT NO. DATE KIND APPLICATION NO. DATE ---------WO 9012591 A1 19901101 WO 1990-CA131 19900426 <--W: AU, CA, JP, KR, SU RW: AT, BE, CH, DE, DK, ES, FR, GB, IT, LU, NL, SE 19940308 19900409 <--US 5292869 Α US 1990-507481 AU 9055261 A1 19901116 AU 1990-55261 19900426 <--AU 649950 B2 19940609 JP 04506794 T2 19921126 JP 1990-506296 19900426 <--JP 3335622 B2 20021021 EP 528787 A1 19930303 EP 1990-906093 19900426 <--EP 528787 В1 19981202 R: AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, LU, NL, SE ES 2127184 Т3 19990416 ES 1990-906093 19900426 <--С 19991214 CA 2051808 CA 1990-2051808 19900426 <--JP 2002316942 A2 20021031 JP 2002-54731 19900426 <--US 6060058 20000509

Α

Α

Α

19890427

19900409 <--

<--

PRAI US 1989-344356

US 1990-507481

US 1995-483881

19950607 <--

```
JP 1990-506296
                          А3
                                19900426 <--
     WO 1990-CA131
                          Α
                                 19900426 <--
     US 1991-639365
                          A3
                                 19910110 <--
                                19920312 <--
     US 1992-851005
                          В1
     US 1994-207719
                          B1
                                19940309
CLASS
                 CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
 WO 9012591
                 ICM
                        A61K039-095
                 ICS
                        A61K039-102; A61K039-02
     {\tt A} method of isolating and purifgying transferrin and lactoferrin receptor
     proteins from bacterial pathogens by affinity chromatog. is described.
     The proteins are used for preparing vaccine antigens. The vaccine antigens
     are effective in preventing diseases caused by bacterial pathogens containing
     lactoferrin and transferrin receptor proteins. The human-lactoferrin
     binding protein from Neisseria meningitidis is identified and
     characterized. It is incorporated into vaccine prepns.
     vaccine bacteria lactoferrin receptor; transferrin receptor vaccine
     bacteria
IT
     Vaccines
        (against bacteria, lactoferrin and transferrin receptors as)
     Receptors
     RL: BIOL (Biological study)
        (for lactoferrin and transferrin, from bacteria, vaccines
        containing)
     Actinobacillus suis
     Haemophilus avium
       Haemophilus gallinarum
       Haemophilus influenzae
       Haemophilus paragallinarum
     Haemophilus pleuropneumoniae
     Haemophilus somnus
       Haemophilus suis
       Moraxella catarrhalis
     Neisseria gonorrhoeae
     Neisseria lactamicus
     Neisseria meningitidis
     Pasteurella haemolytica
     Pasteurella multocida
        (lactoferrin and transferrin receptors from, vaccines containing)
     Lactoferrins
     Transferrins
     RL: BIOL (Biological study)
        (receptors for, from bacteria, vaccines containing)
IT
     Antigens
     RL: BIOL (Biological study)
        (vaccines, against bacteria, lactoferrin and transferrin receptors as)
     Proteins, specific or class
IT
     RL: BIOL (Biological study)
        (lactoferrin-binding, vaccines containing)
IT
     Proteins, specific or class
     RL: BIOL (Biological study)
        (transferrin-binding, vaccines containing)
L44 ANSWER 13 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
ΑN
     1991:171296 HCAPLUS
DN
     114:171296
     Entered STN: 03 May 1991
ED
     Cytokine and hormone carriers for conjugate vaccines
ТT
IN
     Pillai, Subramonia; Eby, Ronald
     Praxis Biologics, Inc., USA
PA
SO
     PCT Int. Appl., 50 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
IC
     ICM A61K047-48
     ICS A61K039-385
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 2, 15
FAN.CNT 2
     PATENT NO.
                         KIND
                                DATE
                                             APPLICATION NO.
                                                                    DATE
                         ----
PΙ
     WO 9101146
                          Al
                                19910207
                                             WO 1990-US3983
                                                                    19900716 <---
         W: AU, CA, FI, JP, KR, NO
         RW: AT, BE, CH, DE, DK, ES, FR, GB, IT, LU, NL, SE
2063271 AA 19910115 CA 1990-2063271
     CA 2063271
                                            CA 1990-2063271
                                                                    19900716 <--
```

```
AU 9060550
                          A1
                                19910222
                                             AU 1990-60550
                                                                    19900716 <--
     AU 651949
                          B2
                                19940811
     EP 482068
                          A1
                                19920429
                                             EP 1990-911070
                                                                    19900716 <--
         R: AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, LU, NL, SE
                                             JP 1990-510469
                                                                    19900716 <--
     JP 04506662
                          T2
                                19921119
     NO 9200160
                          Α
                                19920305
                                             NO 1992-160
                                                                    19920113 <--
PRAI US 1989-380566
                                19890714
                          Α
     WO 1990-US3983
                                19900716
                          Α
CLASS
 PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
                 ICM
                        A61K047-48
WO 9101146
                 ICS
                        A61K039-385
     Immunogenic conjugates are disclosed comprising a carbohydrate-containing
     antigen or other antigen bound to or genetically fused with a cytokine,
     lymphokine, hormone, or growth factor having immunomodulating activity,
     wherein the cytokine, lymphokine, hormone, or growth factor is capable of
     modifying immunogenicity of the carbohydrate-containing antigen. The cytokine
     or lymphokine can be an interleukin or an interferon. The immunogenic
     conjugate can be used in vaccine and covaccine formulations.
ST
     vaccine cytokine hormone carrier
     Animal growth regulators
     Hormones
     Interferons
     RL: PREP (Preparation)
        (antigen bound to, for vaccine preparation)
     Lymphokines and Cytokines
     RL: PREP (Preparation)
        (antigen conjugates, for vaccine preparation)
     Carbohydrates and Sugars, biological studies
     Oligosaccharides
     Polysaccharides, biological studies
     RL: PREP (Preparation)
        (antigens containing, conjugates with cytokines and hormones, for vaccine
IT
     Fungi
     Parasite
     Virus, animal
        (antigens of, conjugates with cytokines and hormones, for vaccine
        preparation)
     Bordetella pertussis
IT
     Clostridium tetani
     Corynebacterium diphtheriae
     Escherichia coli
       Haemophilus influenzae
     Klebsiella pneumoniae
       Moraxella catarrhalis
     Neisseria gonorrhoeae
     Neisseria meningitidis
     Pseudomonas aeruginosa
     Staphylococcus aureus
     Streptococcus pneumoniae
     Streptococcus pyogenes
     Vibrio cholerae
        (capsular polymers of, conjugates with cytokines and hormones, for
        vaccine preparation)
IT
     Antigens
     RL: BIOL (Biological study)
        (carbohydrate and hormone conjugates, in vaccine formulations)
IT
     Vaccines
        (immunogenic conjugates with cytokines and hormones in formulations of)
     Lipopolysaccharides
     RL: PREP (Preparation)
        (of bacteria, conjugates with cytokines and hormones, in vaccine
        preparation)
     Peptidoglycans
     RL: PREP (Preparation)
        (of bacterial cell wall, conjugates with cytokines and hormones, in
        vaccine preparation)
IT
     Lymphokines and Cytokines
     RL: PREP (Preparation)
        (interleukin 1.alpha., antigen bound to, for vaccine preparation)
IT
     Lymphokines and Cytokines
     RL: PREP (Preparation)
        (interleukin 1.beta., antigen bound to, for vaccine preparation)
IT
     Lymphokines and Cytokines
```

```
RL: PREP (Preparation)
        (interleukin 2, antigen bound to, for vaccine preparation)
     Lymphokines and Cytokines
TT
     RL: PREP (Preparation)
        (tumor necrosis factor, antigen conjugates, for vaccine preparation)
IT
     9002-62-4D, Prolactin, antigen conjugates 9002-72-6D, Somatotropin,
     antigen conjugates 9004-10-8D, Insulin, antigen conjugates
     62229-50-9D, Epidermal growth factor, antigen conjugates 62683-29-8D,
     Granulocyte colony-stimulating factor, antigen conjugates 82197-76-0D, Polyribosylribitolphosphate, antigen conjugates 83869-56-1D, Granulocyte
     macrophage colony stimulating factor, antigen conjugates
     RL: BIOL (Biological study)
        (vaccines from)
    ANSWER 14 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
L44
     1990:558671 HCAPLUS
ΑN
     113:158671
DN
ED
     Entered STN: 27 Oct 1990
     T-cell epitope as carrier molecule for conjugate vaccines
ΤI
    Bixler, Garvin; Pillai, Subramonia; Insel, Richard Praxis Biologics, Inc., USA
IN
PA
     PCT Int. Appl., 103 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LΑ
IC
     ICM A61K039-385
     ICS C07K015-04; A61K039-155
CC
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 15
FAN.CNT 1
     PATENT NO.
                         KIND
                                            APPLICATION NO.
                                                                    DATE
                         ----
                          A2
                                19890810
                                            WO 1989-US388
                                                                    19890131 <--
     WO 8906974
     WO 8906974
                          A3
                                19890824
        W: AU, DK, FI, JP, NO
        RW: AT, BE, CH, DE, FR, GB, IT, LU, NL, SE
                                19890825
                                            AU 1989-30654
                                                                    19890131 <--
     AU 8930654
                         A1
     AU 634153
                          B2
                                19930218
     EP 399001
                          A1
                                19901128
                                             EP 1989-908669
                                                                    19890131 <--
     EP 399001
                          B1
                                19940727
        R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE
     JP 03502691
                          T2
                                19910620
                                             JP 1989-502396
                                                                    19890131 <--
     JP 2921574
                          B2
                                19990719
     NO 9002909
                          Α
                                19900827
                                            NO 1990-2909
                                                                    19900629 <--
     NO 179164
                          В
                                19960513
     NO 179164
                          С
                                19960821
     DK 9001829
                          Α
                                19900731
                                            DK 1990-1829
                                                                    19900731 <--
     DK 174416
                          B1
                                20030217
     US 5785973
                          Α
                                19980728
                                            US 1995-481923
                                                                    19950607 <--
PRAI US 1988-150688
                                19880201 <--
                          Α
     US 1989-304783
                          B1
                                19890131 <--
     WO 1989-US388
                          Α
                                19890131 <--
     US 1992-828711
                          B1
                                19920131 <--
                          B1
                                19931209
     US 1993-164989
CLASS
 PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
WO 8906974
                ICM
                        A61K039-385
                        C07K015-04; A61K039-155
                 TCS
US 5785973
                 ECLA A61K039/385; A61K047/48R; C07K014/195
    Conjugates between T-cell epitopes (recognition sites) for bacterial
     products such as tetanus toxin and medically useful substances such as
     antigens, haptens, or antigenic determinants are prepared These conjugates
     elicit antibody responses and are useful in vaccine prepns. The use of
     the T-cell epitope, as opposed to a larger peptide containing the epitope,
     provides an economic advantage in that it may be readily prepared as well as
     a safety advantage in avoidance of use of the whole protein. The
     conjugates also stimulate antibodies against tumor-specific or
     tumor-associated antigens and are useful in the immunization of infants whose
     immune system is not fully developed. The DeLisi and Berzofsky algorithm
     (1985) for potential amphipathic regions was applied to diphtheria toxin
     cross-reactive material (CRM) and 6 regions were identified. Peptides
     corresponding to these CRM regions were synthesized (synthesis given) and
     those stimulating T-cells were conjugated to phosphoribosylribitol
     phosphate (PRP, capsular polymer of Haemophilus influenzae b). The
     conjugates were capable of stimulating antibodies to PRP.
```

```
T lymphocyte epitope conjugate vaccine; safety T lymphocyte epitope
ST
     conjugate vaccine
     Allergy inhibitors
IT
     Neoplasm inhibitors
        (antigen conjugates with T-cell epitopes of bacterial products as)
IT
     Microorganism
     Neoplasm, composition
     Parasite
     Virus
        (antigens of, conjugates with T-cell epitopes of bacterial products, as
        vaccines)
IT
     Haemophilus influenzae
     Neisseria meningitidis
        (capsular antigen and outer membrane protein of, conjugates with T-cell
        epitopes of bacterial products, as vaccines)
IT
     Salmonella typhi
     Streptococcus pneumoniae
        (capsular antigens of, conjugates with T-cell epitopes of bacterial
        products, as vaccines)
IT
     Allergens
       Antigens
     Carbohydrates and Sugars, biological studies
     RL: BIOL (Biological study)
        (conjugates with T-cell epitopes of bacterial products, as vaccines)
     Escherichia coli
       Moraxella catarrhalis
     Neisseria gonorrhoeae
     Streptococcus pyogenes
        (outer membrane proteins of, conjugates with T-cell epitopes of
        bacterial products, as vaccines)
IT
     Lymphocyte
        (B-, epitopes reactive with, conjugates with T-cell epitopes, as
        vaccines)
     Glycoproteins, specific or class
     RL: BIOL (Biological study)
        (F (fusion), conjugates, with T-cell epitopes of bacterial products, as
        vaccine)
\mathbf{T}
     Proteins, specific or class
     RL: BIOL (Biological study)
        (OMP (outer membrane protein), conjugates, with T-cell epitopes of
        bacterial products, as vaccines)
IT
        (T-, bacterial epitope reactive with, conjugates with antigens, as
        vaccines)
IT
    Disease
        (autoimmune, treatment of, antigen conjugates with T-cell epitopes of
        bacterial products for)
IT
     Lipopolysaccharides
     RL: BIOL (Biological study)
        (conjugates, of gram-neg. bacteria, with T-cell epitopes of bacterial
        products, as vaccines)
IT
     Toxins
     Toxoids
     RL: BIOL (Biological study)
        (diphtheria, T-cell epitopes of, antigen conjugates, as vaccines)
        (gram-neg., lipopolysaccharides of, conjugates with T-cell epitopes of
        bacterial products, as vaccines)
IT
     Toxins
     Toxoids
     RL: BIOL (Biological study)
        (pertussis, T-cell epitopes of, antigen conjugates, as vaccines)
IT
     Virus, animal
        (respiratory syncytial, vaccine for, F protein conjugates with T-cell
        epitopes of bacterial products as)
IT
     Toxins
     Toxoids
     RL: BIOL (Biological study)
        (tetanus, T-cell epitopes of, antigen conjugates, as vaccines)
     129774-60-3, Toxin (corynephage .beta. strain ATCC 53281 reduced)
     RL: BIOL (Biological study)
        (T-cell epitopes of, in vaccine preparation)
     128516-95-0D, antigen conjugates
                                         128786-78-7D, antigen conjugates
IT
    129813-87-2D, antigen conjugates 129836-17-5D, antigen conjugates
                                         129813-88-3D, antigen conjugates 129836-18-6D, antigen conjugates
```

```
129851-39-4D, antigen conjugates
     RL: BIOL (Biological study)
        (as vaccines)
                  128786-78-7 129813-87-2 129813-88-3 129836-17-5
     128516-95-0
IT
     129836-18-6
                   129851-39-4
     RL: BIOL (Biological study)
        (in vaccine preparation)
     129813-89-4P 129813-90-7P
                                    129813-91-8P
                                                   129813-92-9P
                                                                   129813-93-0P
TT
     129813-94-1P
                    129813-95-2P
                                    129813-96-3P
                                                   129813-97-4P
                                                                   129813-98-5P
     129813-99-6P
                   129814-00-2P
                                    129814-01-3P
                                                   129836-19-7P
     RL: PRP (Properties); PREP (Preparation)
        (preparation and amino acid sequence of, of diphtheria toxin cross-reactive
        material, in vaccine preparation)
     82197-76-0DP, Polyribosylribitol phosphate, conjugates with synthetic
     peptides of dipHtheria toxin cross-reactive material 129813-89-4DP,
     conjugates with polyribosylribitol phosphate 129813-90-7DP, conjugates
     with polyribosylribitol phosphate 129813-91-8DP, conjugates with
     polyribosylribitol phosphate
                                    129813-92-9DP, conjugates with
     polyribosylribitol phosphate
                                    129813-93-0DP, conjugates with
                                    129813-94-1DP, conjugates with
129813-95-2DP, conjugates with
     polyribosylribitol phosphate
     polyribosylribitol phosphate
     polyribosylribitol phosphate
                                    129813-96-3DP, conjugates with
     polyribosylribitol phosphate polyribosylribitol phosphate
                                     129813-97-4DP, conjugates with
                                    129813-98-5DP, conjugates with
     polyribosylribitol phosphate
                                    129813-99-6DP, conjugates with
     polyribosylribitol phosphate
                                    129814-00-2DP, conjugates with
                                    129814-01-3DP, conjugates with
     polyribosylribitol phosphate
     polyribosylribitol phosphate
                                    129836-19-7DP, conjugates with
     polyribosylribitol phosphate
     RL: PREP (Preparation)
        (preparation of, in preparation of vaccines for Haemophilus influenzae)
L44 ANSWER 15 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
     1990:520772 HCAPLUS
AN
DN
     113:120772
     Entered STN: 29 Sep 1990
ED
    Milk antibody production with shaped polymer microparticles for
TΤ
     controlled-release of antigens
IN
     Beck, Lee R.
PA
     Stolle Research and Development Corp., USA
     U.S., 9 pp. Cont. of U.S. Ser. No. 576,001, abandoned. CODEN: USXXAM
SO
DT
     Patent
LΑ
     English
     ICM A61K039-00
IC
     424088000
NCT.
CC
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 15, 17
FAN.CNT 7
     PATENT NO.
                         KIND
                                DATE
                                             APPLICATION NO.
                                                                     DATE
                          ----
     US 4919929
                          Α
                                19900424
                                             US 1986-910297
                                                                     19860917 <--
                                             AU 1987-76086
     AU 605159
                          B2
                                19910110
                                                                     19870724 <--
                          A1
                                 19890127
     AU 8776086
     IIS 4956349
                                 19900911
                                             US 1988-177223
                                                                     19880404 <--
                          Α
     US 5242691
                          Α
                                19930907
                                             US 1990-580382
                                                                     19900911 <--
     CA 2072658
                          AΑ
                                19910507
                                             CA 1990-2072658
                                                                     19901030 <--
     CA 2072658
                          С
                                20010220
     WO 9106321
                          A1
                                19910516
                                             WO 1990-US6215
                                                                     19901030 <--
         W: AU, CA, JP, NO
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LU, NL, SE
     AU 9066149
                          A1
                                19910531
                                             AU 1990-66149
                                                                     19901030 <--
     AU 644820
                                 19931223
                          B2
     JP 05501801
                          T2
                                 19930408
                                             JP 1990-515068
                                                                     19901030 <--
     JP 2912451
                          B2
                                 19990628
     EP 593440
                          A1
                                 19940427
                                             EP 1990-915892
                                                                     19901030 <--
     EP 593440
                          В1
                                19960821
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE
     AT 141513
                          E
                                19960915
                                             AT 1990-915892
                                                                    19901030 <--
     NO 9201748
                                 19920701
                                             NO 1992-1748
                                                                     19920504 <--
                          Α
     US 5352462
                          Α
                                19941004
                                            US 1992-966741
                                                                    19921027 <--
                                19840201 <--
PRAI US 1984-576001
                          В1
     US 1982-384625
                          B2
                                19820603 <--
     US 1983-576001
                          В1
                                19830201
                                           <--
     US 1983-546162
                          А3
                                 19831027
                                           <--
     US 1986-910297
                                19860917
                          Α
                                           <--
```

```
US 1987-1848
                                19870109 <--
     US 1988-177223
                          A2
                                19880404 <--
     US 1989-431639
                                19891106 <--
                          Α
    US 1990-580382
                          A2
                                19900911 <--
CLASS
 PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
US 4919929
                 ICM
                        A61K039-00
                 NCL
                        424088000
                        A23C009/20; A24B015/30; A61K009/16H6D4; A61K035/20; A61K035/54; A61K035/74; A61K039/395A
US 5352462
                 ECLA
    Milk having elevated IgG antibody levels is produced by (1) i.m. or s.c.
     implantation within a bovidae of a hyperimmunization response-eliciting
     amount of an antigenic substance incorporated within shaped microparticles
     of a biocompatible matrix material which causes controlled-release of the
     antigen, thereby prolonging antigenic activity within the bovidae; and (2)
     recovering milk having an elevated level of antibody. The immunized state
     may be attained more rapidly by simultaneously administering the shaped
    matrix and the antigenic substance in liquid form. A polyvalent antigen
     sample (S-100) containing .apprx.26 kinds of heat-killed bacteria was
     microencapsulated in biodegradable lactide-glycolide copolymer
     microparticles (diameter < 250 .mu.m). The microparticles were suspended in
     vehicle (Tween 20 and CMC) and injected i.m. one time into cows.
     Increased antibody titer was shown in the milk.
st
     controlled release antigen vaccine; cattle milk antibody prodn
IT
     Vaccines
        (antigens in controlled-release microparticles as, for elevated milk
        antibody production)
IT
    Actinobacillus equuli
     Actinobacillus lignieresii
     Actinobacillus seminis
     Bacillus cereus
     Brucella melitensis
     Campylobacter fetus
     Campylobacter fetus intestinalis
     Cell
     Chlamydia psittaci
     Clostridium tetani
     Corynebacterium pyogenes
     Corynebacterium renale
     Enterobacter aerogenes
     Escherichia coli
     Fusobacterium necrophorum
     Gardnerella vaginalis
      Haemophilus ducreyi
       Haemophilus influenzae
     Klebsiella pneumoniae, oxides
     Leptospira interrogans pomona
     Listeria monocytogenes
      Moraxella bovis
     Mycobacterium tuberculosis
     Mycoplasma bovigenitalium
     Mycoplasma hominis
     Neisseria gonorrhoeae
     Pasteurella haemolytica
     Pasteurella multocida
     Propionibacterium acnes
     Proteus vulgaris
     Pseudomonas aeruginosa
     Pseudomonas maltophilia
     Rhodococcus equi
     Salmonella abortivoequina
     Salmonella abortusovis
     Salmonella dublin
     Salmonella enteritidis
     Salmonella heidelberg
     Salmonella paratyphi
     Salmonella typhimurium
     Shiqella dysenteriae
     Staphylococcus aureus
     Staphylococcus epidermidis
     Streptococcus agalactiae
     Streptococcus bovis
     Streptococcus dysgalactiae
     Streptococcus equisimilis
```

Streptococcus mitis

```
Streptococcus mutans
     Streptococcus pneumoniae
     Streptococcus pyogenes
     Streptococcus salivarius
     Streptococcus sanguis
     Streptococcus uberis
    Treponema pallidum
     Virus, animal
     Yersinia enterocolitica
        (antigens of, polymer microparticles containing, for controlled-release
        vaccines and elevated milk antibody production)
IT
    Milk
        (elevated antibody production in, controlled-release microencapsulated
       antigen vaccines for)
    Disease
TT
        (lymphopathia venereum, antigens of, polymer microparticles containing, for
        controlled-release vaccines and elevated milk antibody production)
IT
    Antibodies
     RL: PRP (Properties)
        (milk containing elevated levels of, production of, controlled-release
        microencapsulated antigen vaccine for)
TТ
     Antigens
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (polymer microparticles containing, for controlled-release vaccines and
        elevated milk antibody production)
TT
    Cattle
        (vaccination of, with controlled-release microencapsulated antigen
        vaccines)
    Bovidae
        (vaccination of, with controlled-release microencapsulated antigen
        vaccines, for production of milk containing elevated levels of IgG antibodies)
TT
    Immunoglobulins
     RL: BSU (Biological study, unclassified); MFM (Metabolic formation); BIOL
     (Biological study); FORM (Formation, nonpreparative)
        (G, milk containing elevated levels of, production of, controlled-release
       microencapsulated antigen vaccine for)
ΙT
    Venereal disease
        (granuloma inguinale, antigens of, polymer microparticles containing, for
        controlled-release vaccines and elevated milk antibody production)
IT
    Streptococcus
        (group B, antigens of, polymer microparticles containing, for
        controlled-release vaccines and elevated milk antibody production)
    Polyethers, biological studies
IT
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (ortho esters, antigen microencapsulation in, for controlled-release
        vaccines and elevated milk antibody production)
                 25248-42-4, Poly[oxy(1-oxo-1,6-hexanediyl)]
IT
    24980-41-4
                                                                25266-42-6
     26009-03-0, Poly[oxy(1-oxo-1,2-ethanediyl)] 26023-30-3
                                               26100-51-6
     Poly[oxy(1-methyl-2-oxo-1,2-ethanediyl)]
                                                            26124-68-5
     31621-87-1
                34346-01-5
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (antigen microencapsulation in, for controlled-release vaccines and
        elevated milk antibody production)
   ANSWER 16 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
L44
    1990:42564 HCAPLUS
ΑN
       Correction of: 1988:461449
DN
    112:42564
       Correction of: 109:61449
    Entered STN: 04 Feb 1990
ED
    Complexes of vitamin B12 and biologically active agents for oral drug
ΤI
    delivery
    Russell-Jones, Gregory John; De Aizpurua, Henry James; Howe, Peter Allan;
IN
    Burge, Geoffery Lewis
PA
    Biotechnology Australia Pty. Ltd., Australia
    Eur. Pat. Appl., 19 pp.
    CODEN: EPXXDW
DT
    Patent
    English
LΑ
IC
    ICM A61K047-00
    ICS A61K037-02; A61K037-24
CC
    63-5 (Pharmaceuticals)
    Section cross-reference(s): 1
```

```
FAN.CNT 1
    PATENT NO.
                       KIND DATE
                                          APPLICATION NO.
                                                                DATE
                              -----
     _____
                       ----
    EP 220030
                        A2
                              19870429
                                          EP 1986-307849
                                                                19861010 <--
                              19870930
    EP 220030
                        A3
    EP 220030
                         B1
                              19910619
        R: AT, BE, CH, DE, ES, FR, GB, GR, IT, LI, LU, NL, SE
                                          ZA 1986-7703
    ZA 8607703
                A
                              19870624
                                                                19861009 <--
    IN 165029
                        Α
                              19890805
                                          IN 1986-CA738
                                                                19861009 <--
    IL 80264
                        A1
                              19911121
                                          IL 1986-80264
                                                                19861009 <--
                                          CA 1986-520162
    CA 1330791
                              19940719
                                                                19861009 <--
                        A1
    WO 8702251
                        A1
                              19870423
                                          WO 1986-AU299
                                                                19861010 <--
        W: AU, DK, FI, JP, KR, NO, SU, US
                A1
                            19870505
    AU 8665289
                                          AU 1986-65289
                                                                19861010 <--
    AU 587658
                        B2
                              19890824
    CN 86107590
                        Α
                              19870520
                                          CN 1986-107590
                                                                19861010 <--
    CN 1045391
                        В
                              19991006
    JP 63501015
                         T2
                              19880414
                                          JP 1986-505525
                                                                19861010 <--
    JP 08000779
                              19960110
                        B4
                                          AT 1986-307849
    AT 64534
                        E
                              19910715
                                                                19861010 <--
    ES 2051690
                        Т3
                              19940701
                                          ES 1986-307849
                                                                19861010 <--
    DK 8702925
                        Α
                              19870609
                                          DK 1987-2925
                                                                19870609 <--
                        В1
    DK 167099
                              19930830
    US 5428023
                              19950627
                                          US 1993-61343
                                                                19930517 <--
                         Α
    US 5589463
                        Α
                              19961231
                                          US 1995-479635
                                                                19950607 <--
                                         US 1995-483811
    US 5807832
                         Α
                              19980915
                                                                19950607 <--
PRAI AU 1985-2838
                        Α
                              19851010 <--
    EP 1986-307849
                              19861010 <--
                        Α
    WO 1986-AU299
                        Α
                              19861010 <--
                                        <--
    US 1987-84821
                        B1
                              19870609
    AU 1988-2838
                        Α
                              19881010 <--
    US 1990-600137
                              19901019 <--
                        B1
    US 1991-759697
                        B1
                              19910909 <--
    US 1993-61343
                        A3
                              19930517
CLASS
               CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
               _ - - - -
                       ______
 EP 220030
               ICM A61K047-00
                ICS
                       A61K037-02; A61K037-24
                     A61K039/00; A61K039/35; A61K039/385; A61K047/48H4
US 5589463
                ECLA
                      A61K047/48H4
US 5807832
               ECLA
    Vitamin B12 is covalently bonded to biol. active substances such as
    hormones, proteins, antigens, haptens, and antibiotics. The B12 in the
    complex can still react with intrinsic factor in the intestine, so the
    natural uptake mechanism for B12 is utilized to deliver various otherwise
    nonabsorbable compds. to the circulation. Vitamin B12 was hydrolyzed to
    give the monocarboxylic acid, which was coupled to N-hydroxysuccinamide,
    and treated with Lys-6-leutinizing hormone releasing hormone (I) to give
    B12-I. This complex induced ovulation in mice following oral
    administration, whereas LHRH, orally, did not induce ovulation.
    B12-neomycin complex was as effective orally in mice against Salmonella
    typhimurium as i.m. neomycin or i.m. B12-neomycin, whereas neomycin orally
    was ineffective.
    oral absorption drug vitamin B12 complex; intestine absorption drug
ST
    vitamin B12 complex; vaccine vitamin B12 complex oral absorption; hormone
    vitamin B12 complex oral absorption; antibiotic vitamin B12 complex oral
    absorption
    Blood-brain barrier
IT
    Intestine, metabolism
       (absorption by, of vitamin B12-biol. active agent complexes)
       (antigen of, complexes with vitamin B12, for oral administration to
       stimulate immune response)
IT
    Kapok
       (antigens of, complexes with vitamin B12, for oral administration to
       stimulate immune response)
    Wheat
IT
       (chaff, antigens of, complexes with vitamin B12, for oral
       administration to stimulate immune response)
IT
    Antibiotics
       (complexes with vitamin B12, for oral administration)
IT
    Allergens
      Haptens
    Hormones
    RL: BIOL (Biological study)
       (complexes with vitamin B12, for oral delivery)
```

```
Albumins, compounds
     Interferons
     RL: BIOL (Biological study)
        (complexes with vitamin B12, for oral drug delivery)
     Dermatophagoides pteronyssinus
     Felidae
     Swine
        (hair of, antigen of, complexes with vitamin B12, for oral
        administration to stimulate immune response)
IT
    Placenta
        (passage of substances across, vitamin B12-active agent complexes for)
IT
     Cholera
     Coccidiosis
     Diphtheria
       Haemophilus influenzae
     Influenza
     Klebsiella pneumoniae
     Measles
       Moraxella catarrhalis
     Mycobacterium BCG
     Plaque
     Rubella
     Salmonella typhi
     Streptococcus
     Streptococcus pneumoniae
     Tetanus
     Tuberculosis
     Variola
     Yellow fever
        (protein derived from or immunogens against, complexes with vitamin B12
        for oral vaccination)
IT
     Ovulation
        (stimulation of, by orally administered vitamin B12-LHRH complex)
     Allergy inhibitors
IT
        (vitamin B12 complexes with hapten or antigen for)
     Immunomodulators
IT
        (vitamin B12-biol. active agent complexes as)
TТ
     Antigens
     RL: BIOL (Biological study)
        (B12, complexes with vitamin, for oral delivery)
     Proteins, specific or class
     RL: BIOL (Biological study)
        (complexes, with vitamin B12, for oral delivery)
IT
     Polysaccharides, compounds
     RL: BIOL (Biological study)
        (complexes, with vitamin B12, for oral drug delivery)
IT
     Embryo
        (fetus, drug delivery in, vitamin B12-active agent complexes for)
IT
     Vaccines
        (oral, vitamin B12 complexes with antigens as)
                 37434-06-3 38285-78-8
                                                           57757-57-0
                                                                         62069-75-4
IT
     6539-14-6
                                            57683-72-4
     68181-17-9
                  74676-97-4
                                74676-98-5
                                             81069-02-5
                                                            98897-08-6
                   111105-75-0
                                  111105-76-1
     102568-45-6
     RL: BIOL (Biological study)
        (coupling agent, for preparation of vitamin B12-biol. active agent
        complexes)
IT
     6066-82-6P
     RL: PREP (Preparation)
        (crosslinking agent for preparation of vitamin B12-biol. active agent
        complexes)
     68-19-9DP, Vitamin B12, complexes with bovine serum albumin
TT
     RL: PREP (Preparation)
        (preparation and stimulation of immune response by, in oral administration)
     88326-63-0DP, Zincobinamide, complexes with biol. active agents
IT
     111070-88-3DP, complexes with biol. active agents
     RL: PREP (Preparation)
        (preparation of, for oral drug administration)
     51-17-2DP, 1H-Benzimidazole, derivs., complexes with biol. active agents
                                          58-14-0DP, vitamin B12 complexes 59-47-2DP, vitamin B12 complexes
     57-42-1DP, vitamin B12 complexes 58-22-0DP, vitamin B12 complexes
     61-32-5DP, vitamin B12 complexes
                                          61-33-6DP, vitamin B12 complexes
     68-19-9DP, Vitamin Bl2, complexes with biol. active agents
                                                                     1404-04-2DP,
     Neomycin, vitamin B12 complexes 1867-66-9DP, vitamin B12 complexes
                                          7361-61-7DP, vitamin B12 complexes 9002-70-4DP, vitamin B12 complexes
     4697-36-3DP, vitamin Bl2 complexes
     9002-61-3DP, vitamin Bl2 complexes
```

```
9004-10-8DP, Insulin, vitamin Bl2 complexes
                                                     9004-66-4DP, Iron dextran,
     vitamin B12 complexes 9034-40-6DP, Luteinizing hormone-releasing factor, vitamin B12 complexes 13408-75-8DP, complexes with biol. active agents
     13422-51-0DP, complexes with biol. active agents 13422-52-1DP, complexes
     with biol. active agents 13422-55-4DP, complexes with biol. active
     agents 13870-90-1DP, complexes with biol. active agents 14978-39-3DP,
     complexes with biol. active agents 15041-07-3DP, complexes with biol.
     active agents 15671-27-9DP, complexes with biol. active agents
     18559-94-9DP, vitamin B12 complexes 20623-13-6DP, complexes with biol.
     active agents 23208-66-4DP, complexes with biol. active agents 23388-02-5DP, complexes with biol. active agents 52671-12-2DP, vitamin
     B12 complexes 57285-09-3DP, Inhibin, vitamin B12 complexes
     112076-75-2DP, complexes with biol. active agents
     RL: PREP (Preparation)
        (preparation of, for oral drug delivery)
L44 ANSWER 17 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN
     1981:145332 HCAPLUS
AN
     94:145332
DN
    Entered STN: 12 May 1984
ED
     Antigens and vaccines containing them
TI
    Hours, Michel; Pourquier, Andre
IN
PA
     Fr.
     Fr. Demande, 11 pp.
SO
     CODEN: FRXXBL
DT
     Patent
LΑ
    French
    A61K039-02; C12K005-00
TC
CC
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 15
FAN.CNT 1
                         KIND DATE
                                             APPLICATION NO.
     PATENT NO.
                                                                      DATE
                                 -----
                          _ _ _ _
     FR 2446111
                          A1
                                 19800808
                                              FR 1979-1301
                                                                      19790112 <--
     FR 2446111
                           B1
                                 19820702
PRAI FR 1979-1301
                                 19790112 <--
                          Α
CLASS
PATENT NO.
                CLASS PATENT FAMILY CLASSIFICATION CODES
FR 2446111
                IC
                        A61K039-02IC
                                          C12K005-00
AB Antigenic complexes were isolated from organisms (Streptococcus pyogenes,
     S. aureus, Diplococcus pneumoniae, Neisseria catarrhalis, N. elongata,
     Escherichia coli, Klebsiella pneumoniae, Hemophilus influenzae, Proteus
     mirabilis, Pseudomonas aeruginosa) by an improved process in which the
     cell membrane was isolated and then solubilized by the simultaneous action
     of Na deoxycholate [302-95-4] (0.5-5 mg/mg proteins), lysozyme [9001-63-2] (1 mg/mL), and EDTA Na [64-02-8] (25 mM) at pH .apprx.7 for
     12-24 h at 20.degree.. The extract was centrifuged and the supernatant liquid
     could be used directly as a source of antigens without removal of the
     solubilizers. The solubilizers protected the antigenic complexes against
     contaminant proteases. The preparation of vaccines from these antigens is
     discussed.
   antigen bacteria vaccine
    Branhamella catarrhalis
     Escherichia coli
       Haemophilus influenzae
     Klebsiella pneumoniae
     Neisseria elongata
     Proteus mirabilis
     Pseudomonas aeruginosa
     Staphylococcus aureus
     Streptococcus pneumoniae
     Streptococcus pyogenes
        (antigen separation from cell membranes of, for vaccine manufacture)
ΙT
     Vaccines
       (manufacture of, antigen separation for)
IТ
     Antigens
     RL: PROC (Process)
        (of bacteria cell membranes, separation of, for vaccines)
IT
     302-95-4
     RL: BIOL (Biological study)
        (bacteria cell membrane solubilization by EDTA and lysozyme and, in
        antigen preparation)
IT
     9001-63-2
     RL: BIOL (Biological study)
        (bacteria cell membrane solubilization by deoxycholate and EDTA and, in
```

```
antigen preparation)
IT
     64-02-8
     RL: BIOL (Biological study)
         (bacteria cell membrane solubilization by deoxycholate and lysozyme
        and, in antigen preparation)
L44 ANSWER 18 OF 18 HCAPLUS COPYRIGHT 2004 ACS on STN AN 1973\!:\!470125 HCAPLUS
     79:70125
     Entered STN: 12 May 1984
Autogenous vaccine. Preparation technique and efficiency factors
ED
ΤI
ΑU
     Cury, Rolando
CS
     Fac. Med. Vet. Zootec., Univ. Sao Paulo, Sao Paulo, Brazil
     Revista de Saude Publica (1972), 6(4), 371-83
so
     CODEN: RSPUB9; ISSN: 0034-8910
DT
     Journal
     Portuguese
LА
     63-3 (Pharmaceuticals)
CC
     Seeding procedures were used to obtain antigenous vaccines. Bacteria were
AB
     preselected. Media were tested for sterility before seeding by heating
     them at 37.degree. for 24 hr. Simple media (pH 7.4), agar (pH 7.4),
     semisolid agar (pH 6.8-7.0), thioglycolate-dextrose, and agar-triptose
     were used. Cultures of Streptococcus, Staphylococcus, Enterobacteriaceae,
     Haemophilus, Bordetella, Pasteurella, Moraxella, Clostridium, Pseudomonas,
     Neisseria, and Brucella were prepared Adequate culture conditions are given in each case. I2, HCHO, and the heat were used to inactivate the bacteria
     to prevent damage of existing antigenous agents. Instructions to apply
     vaccines prepared are given.
ST
     bacteria vaccine
IT
    Bacteria
     Bordetella
     Brucella
     Clostridium
     Enterobacteriaceae
       Haemophilus
       Moraxella
     Neisseria
     Pasteurella
     Pseudomonas
     Staphylococcus
     Streptococcus
        (antigens of, vaccines of)
IT
     Vaccines
        (of bacteria antigens)
IT
     Antigens
     RL: BIOL (Biological study)
        (of bacteria, vaccines of)
=> d all 136 tot
L36 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2004 ACS on STN
     2000:741941 HCAPLUS
     133:320987
DN
     Entered STN: 20 Oct 2000
ED
TI
     Conserved adhesin motif and methods of use thereof
IN
     Lupas, Andrei Nicolae
PΑ
     Smithkline Beecham Corporation, USA; Smithkline Beecham PLC
     PCT Int. Appl., 85 pp.
SO
     CODEN: PIXXD2
חת
     Patent
LA 'English
IC
     ICM A61K038-00
          A61K039-00; A61K039-395; C07H021-04; C07K001-00; C07K016-00;
          C12N005-00; C12N007-00; C12N015-09; C12P021-08; G01N033-53
     15-2 (Immunochemistry)
     Section cross-reference(s): 1, 3, 63
FAN.CNT 1
     PATENT NO.
                          KIND
                                  DATE
                                               APPLICATION NO.
                                                                       DATE
     WO 2000061165
                           A1
                                  20001019
                                               WO 2000-US9866
                                                                       20000413 <--
         W: JP, US
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE
PRAI US 1999-129073P
                                  19990413 <--
CLASS
```

Graser 10/030313 Page 42

```
PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
 WO 2000061165
                        A61K038-00
                 ICM
                        A61K039-00; A61K039-395; C07H021-04; C07K001-00;
                 ICS
                        C07K016-00; C12N005-00; C12N007-00; C12N015-09;
                        C12P021-08; G01N033-53
     Isolated polypeptides which are conserved in eubacterial extracellular
AB
     domains are identified in five pathogens of the beta and gamma branches of
     proteobacteria. These polypeptides, alone or as fusion proteins with a
     second protein, are useful in the generation of antibodies or other
     antagonists. The peptides, fusion proteins, and antibodies are useful as
     vaccine components or therapeutic agents against bacterial infection or as
     diagnostic reagents. These polypeptides are also useful in screening
     methods for other agonists and antagonists which may be used in diagnosis,
     therapy, and as vaccines.
     proteobacteria infection adhesin motif antibody vaccine
ST
     Chaperonins
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (DnaK, fusion protein; conserved adhesin motif and fusion proteins for
        use as vaccine or diagnostic agent and therapeutic agent against
        bacterial infection)
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (GST, fusion protein; conserved adhesin motif and fusion proteins for
        use as vaccine or diagnostic agent and therapeutic agent against
        bacterial infection)
ΙT
    Heat-shock proteins
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (HSP 70, fusion protein; conserved adhesin motif and fusion proteins
        for use as vaccine or diagnostic agent and therapeutic agent against
        bacterial infection)
TΤ
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (NS1 (nonstructural, 1), fusion protein; conserved adhesin motif and
        fusion proteins for use as vaccine or diagnostic agent and therapeutic
        agent against bacterial infection)
     Proteins, specific or class
IT
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (UspAl; conserved adhesin motif and fusion proteins for use as vaccine
        or diagnostic agent and therapeutic agent against bacterial infection)
     Proteins, specific or class
TT
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (UspA2; conserved adhesin motif and fusion proteins for use as vaccine
        or diagnostic agent and therapeutic agent against bacterial infection)
     Proteins, specific or class
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (YadA outer membrane adhesin; conserved adhesin motif and fusion proteins for use as vaccine or diagnostic agent and therapeutic agent
        against bacterial infection)
TΤ
     Receptors
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (adhesin; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
IT
     Immunostimulants
        (adjuvants; conserved adhesin motif and fusion proteins for
        use as vaccine or diagnostic agent and therapeutic agent against
        bacterial infection)
IT
     Antibodies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (anti-idiotypic; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
IT
     Infection
        (bacterial; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
```

IT

Drug delivery systems

```
(carriers; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
TT
     Antigens
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (composition; conserved adhesin motif and fusion proteins for use as vaccine
        or diagnostic agent and therapeutic agent against bacterial infection)
    Actinobacillus
     Bacteria (Eubacteria)
     Drug screening
     Escherichia coli
       Haemophilus
     Influenza
     Labels
       Moraxella
     Mycobacterium
     Neisseria
     Pathogen
     Protein motifs
     Protein sequences
     Proteobacteria
     Simulation and Modeling, physicochemical
       Vaccines
     Yersinia
     Yersinia pestis
        (conserved adhesin motif and fusion proteins for use as vaccine or
        diagnostic agent and therapeutic agent against bacterial infection)
     Fusion proteins (chimeric proteins)
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (conserved adhesin motif and fusion proteins for use as vaccine or
        diagnostic agent and therapeutic agent against bacterial infection)
IT
     Adhesins
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
    Antibodies
     Nucleic acids
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (conserved adhesin motif and fusion proteins for use as vaccine or
        diagnostic agent and therapeutic agent against bacterial infection)
тт
     Toxoids
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (diphtheria, fusion protein; conserved adhesin motif and fusion
        proteins for use as vaccine or diagnostic agent and therapeutic agent
        against bacterial infection)
IT
     Immunoglobulins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (fragments; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
     .alpha.-Factor (microbial)
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (fusion protein; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
IT
    Diagnosis
        (immunodiagnosis; conserved adhesin motif and fusion proteins for use
        as vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
IT
    Animal cell
        (mammalian; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
IT
    Antibodies
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (monoclonal; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
```

```
infection)
IT
     Animal virus
        (recombinant; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
     Genetic element
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (regulatory sequence; conserved adhesin motif and fusion proteins for
        use as vaccine or diagnostic agent and therapeutic agent against
        bacterial infection)
TΤ
     Toxoids
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (tetanus, fusion protein; conserved adhesin motif and fusion proteins
        for use as vaccine or diagnostic agent and therapeutic agent against
        bacterial infection)
     302579-96-0
IT
                  302579-98-2
     RL: PRP (Properties)
        (Unclaimed; conserved adhesin motif and methods of use thereof)
ΤТ
     301857-38-5
                   302323-37-1
                                 302323-38-2
                                                302323-39-3
                                                              302323-42-8
                   302323-49-5
                                 302323-50-8
                                                302323-51-9
                                                              302323-52-0
     302323-48-4
     302323-53-1
                   302323-54-2
                                 302323-55-3
                                                302323-56-4
                                                              302323-57-5
     302352-24-5
                   302798-58-9
                                302798-59-0
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (conserved adhesin motif and fusion proteins for use as vaccine or
        diagnostic agent and therapeutic agent against bacterial infection)
     60267-61-0P, Ubiquitin
TΤ
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (fusion protein; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
IT
     9030-53-9. Galactokinase
                               9031-11-2, .beta.-Galactosidase
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (fusion protein; conserved adhesin motif and fusion proteins for use as
        vaccine or diagnostic agent and therapeutic agent against bacterial
        infection)
TT
     302684-47-5
                   302684-48-6
                                 302684-49-7
                                                302684-50-0
                                                              302684-51-1
     302684-52-2
                   302684-53-3
                                 302684-54-4
                                                302684-55-5
                                                              302684-56-6
     302684-57-7
                   302684-58-8
                                 302684-59-9
                                                302684-60-2
                                                              302684-61-3
     302684-62-4
                   302684-63-5
                                 302684-64-6
                                                302684-65-7
                                                              302684-66-8
                   302684-68-0
                                 302684-69-1
                                                302684-70-4
                                                              302684-71-5
     302684-67-9
     302684-72-6
                   302684-73-7
                                 302684-74-8
                                                302684-75-9
                                                              302684-76-0
     302684-77-1
                   302684-78-2
                                 302684-79-3
                                                302684-80-6
                                                              302684-81-7
     302684-82-8
                   302684-83-9
                                 302684-84-0
                                                302684-85-1
                                                              302684-86-2
     302684-87-3
                   302684-88-4
                                 302684-89-5
                                                302684-90-8
                                                              302905-45-9
                   302905-52-8
                                 302905-61-9
                                                302905-74-4
                                                              302906-00-9
     302905-47-1
     302906-01-0
                   302906-02-1
     RL: PRP (Properties)
        (unclaimed protein sequence; conserved adhesin motif and methods of use
        thereof)
    302323-40-6
                   302323-41-7
                                 302323-43-9
                                                302323-44-0
                                                              302323-45-1
     302323-46-2
                   302323-47-3
                                 302323-58-6
                                                302323-59-7
                                                              302323-60-0
     302323-61-1
                   302323-62-2
                                 302323-63-3
                                                302323-64-4
                                                              302323-65-5
     302323-66-6
                   302325-08-2
                                 302325-09-3
                                                302325-10-6
                                                              302325-11-7
                                 302579-65-3
                                                302579-67-5
     302579-61-9
                   302579-63-1
                                                              302579-69-7
     302579-71-1
                   302579-74-4
                                 302579-81-3
                                                302579-86-8
                                                              302579-88-0
     302579-91-5
                   302579-99-3
                                 302580-00-3
                                                302580-01-4
                                                              302580-02-5
     302580-04-7
                   302580-05-8
                                 302580-06-9
                                                302580-07-0
                                                              302580-08-1
                                 302580-11-6
                                                302580-12-7
     302580-09-2
                   302580-10-5
                                                              302580-13-8
     302580-16-1
                   302580-19-4
                                 302580-20-7
                                                302580-22-9
                                                              302580-24-1
     302580-25-2
                   302580-26-3
                                 302580-27-4
                                                302580-28-5
                                                              302580-29-6
     302580-30-9
                   302580-31-0
                                 302580-32-1
                                                302580-33-2
                                                              302580-34-3
     302580-35-4
                   302580-36-5
                                 302580-38-7
                                                302580-39-8
                                                              302580-40-1
     302580-41-2
                   302580-42-3
                                 302684-46-4
                                                302798-57-8
     RL: PRP (Properties)
        (unclaimed sequence; conserved adhesin motif and methods of use
        thereof)
RE.CNT
              THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
(1) Skurnik; Molecular Microbiology 1989, V3, P517 HCAPLUS
```

The Board Of Regents The University Of Texas System; WO 9828333 A2 1998

**HCAPLUS** 

```
=> d all 142 tot
     ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
L42
     1999:495189 HCAPLUS
ΆN
DN
     131:129044
     Entered STN: 10 Aug 1999
ED
     Vaccine composition comprising milled lyophilizate of antigenic whole
     cells
IN
     Hafner, Roderick Peter
PA
     Raby Limited, UK
     PCT Int. Appl., 33 pp.
     CODEN: PIXXD2
     Patent
דת
     English
LΑ
     ICM A61K039-00
IC
     ICS A61K039-02
     15-2 (Immunochemistry)
CC
     Section cross-reference(s): 63
FAN.CNT 1
     PATENT NO.
                          KIND
                                 DATE
                                              APPLICATION NO.
                                                                      DATE
                          ----
                                 19990805
                                              WO 1999-GB287
DT
     WO 9938529
                           A1
                                                                      19990128 <--
         W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP,
             KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,
             MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
             CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     AU 9922897
                           A1
                                 19990816
                                            AU 1999-22897
                                                                      19990128 <--
PRAI GB 1998-1870
                           Α
                                 19980128
     WO 1999-GB287
                                 19990128 <--
                           W
CLASS
 PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
                         A61K039-00
 WO 9938529
                 ICM
                         A61K039-02
                 ICS
WO 9938529
                 ECLA
                         A61K039/00; A61K039/102; A61K039/02
    A vaccine composition for the prevention of bacterial or fungal infections of
     mucosal surfaces comprises a lyophilisate of antigenic whole cells milled
     to a particle size of from about 20 to 350<mm. The vaccine may contain
     killed organisms such as Haemophilus influenzae or Pseudomonas aeruginosa
     and is useful, for example, for preventing the colonization by these
     organisms of patients suffering from chronic lung diseases or, in
     previously colonized patients, for preventing the occurrence of acute
     infection of the respiratory tract.
ST
     vaccine lyophilized bacteria fungus yeast antigen
IT
     Immunostimulants
        (adjuvants; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
TT
     Infection
        (bacterial, secondary; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
     Drug delivery systems
IT
        (capsules; vaccine composition comprising milled lyophilizate of antigenic
        whole cells)
IT
     Eye, disease
        (conjunctivitis, bacterial; vaccine composition comprising milled
        lyophilizate of antigenic whole cells)
TT
     Ear
        (disease, eustachian tube infection; vaccine composition comprising milled
        lyophilizate of antigenic whole cells)
IT
     Respiratory tract
        (disease, exacerbation; vaccine composition comprising milled lyophilizate
        of antigenic whole cells)
IT
     Mammary gland
     Urogenital tract
        (disease, infection; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
IT
        (diseases, infection; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
```

Page 46

```
IT
    Escherichia coli
        (enteropathogenic; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
    Escherichia coli
IT
        (enterotoxigenic; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
IT
    Digestive tract
        (gastroenteritis; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
IT
    Digestive tract
    Eye, disease
    Mouth
    Respiratory tract
    Urogenital tract
    Vagina
        (infection; vaccine composition comprising milled lyophilizate of antigenic
IT
    Ear
        (middle, infection; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
ΙT
    Pharynx
        (nasopharynx, infection; vaccine composition comprising milled lyophilizate
       of antigenic whole cells)
IT
    Ear
        (otitis, otitis media; vaccine composition
        comprising milled lyophilizate of antigenic whole cells)
ΙT
    Pharynx
        (pharyngitis; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
    Drug delivery systems
IT
        (powders; vaccine composition comprising milled lyophilizate of antigenic
       whole cells)
IT
    Drug delivery systems
        (tablets; vaccine composition comprising milled lyophilizate of antigenic
        whole cells)
IT
    Tonsil
        (tonsillitis; vaccine composition comprising milled lyophilizate of
        antigenic whole cells)
IT
    Digestive tract
        (ulcer; vaccine composition comprising milled lyophilizate of antigenic
       whole cells)
IT
    Bacteria (Eubacteria)
    Burkholderia cepacia
    Candida
    Candida albicans
    Candida glabrata
    Candida krusei
    Chlamydia trachomatis
    Cholera
    Common cold
    Corynebacterium parvum
    Diarrhea
    Diphtheria
    Fermentation
    Fungi
    Granulicatella adiacens
      Haemophilus influenzae
    Helicobacter pylori
    Klebsiella pneumoniae
    Klebsiella pneumoniae ozaenae
    Lactococcus lactis
    Meningitis
    Microorganism
      Moraxella catarrhalis
    Mycobacterium BCG
    Mycobacterium tuberculosis
    Mycosis
    Neisseria gonorrhoeae
    Neisseria meningitidis
    Pertussis
    Pneumonia
    Pseudomonas
    Pseudomonas aeruginosa
    Salmonella typhi
    Sexually transmitted diseases
```

Staphylococcus aureus

```
Streptococcus
     Streptococcus mutans
     Streptococcus pneumoniae
     Streptococcus pyogenes
     Tuberculosis
     Typhoid fever
       Vaccines
     Vibrio cholerae
     Virus
     Yeast
        (vaccine composition comprising milled lyophilizate of antigenic whole
        cells)
IT
     Antigens
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (vaccine composition comprising milled lyophilizate of antigenic whole
              THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Angus, R; DEVELOPMENTS IN BIOLOGICAL STANDARDIZATION 1984, V56, P659
    MEDLINE
(2) Esquisabel, A; JOURNAL OF MICROENCAPSULATION 1997, V14(5), P627 HCAPLUS
     ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
1.42
     1999:464181 HCAPLUS-
AN
DN
     131:86860
     Entered STN: 29 Jul 1999
Lipooligosaccharide-based accine for prevention of Moraxella
ED
ТT
     (Branhamella) catarrhalis infections in mammals
IN
     Gu, Xin-Xing; Robbins, John B.
PA
     The Government of the United States of America, Department of Health and
     Human, USA
SO
     PCT Int. Appl., 60 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
     ICM A61K039-02
IC
     ICS A61K039-385; C08B037-00
     15-2 (Immunochemistry)
CC
     Section cross-reference(s): 63
FAN. CNT 3
     PATENT NO.
                          KIND
                                 DATE
                                              APPLICATION NO.
                                                                      DATE
     WO 9936086
                                 19990722
                                              WO 1999-US590
                                                                      19990112 <--
PΤ
                           A1
         W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,
             MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
             TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
             FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
             CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     CA 2315746
                           AA
                                 19990722
                                              CA 1999-2315746
                                                                      19990112 <--
     AU 9922212
                           A1
                                 19990802
                                              AU 1999-22212
                                                                      19990112 <--
     BR 9906902
                           Α
                                 20001017
                                              BR 1999-6902
                                                                       19990112 <--
     EP 1047447
                           A1
                                 20001102
                                              EP 1999-902170
                                                                      19990112 <--
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
     JP 2002509115
                           T2
                                 20020326
                                              JP 2000-539859
                                                                       19990112 <--
     US 6685949
                                 20040203
                                              US 2000-610034
                                                                      20000705 <--
                           В1
     US 2004126381
                           A1
                                 20040701
                                              US 2003-688115
                                                                      20031017 <--
                                              US 2003-729027
     US 2004115214
                           A1
                                 20040617
                                                                      20031205 <--
PRAI US 1998-71483P
                           P
                                 19980113
                                            <--
     US 1996-16020P
                           Ρ
                                 19960423
                                            <--
                                            <--
     US 1997-842409
                           А3
                                 19970423
     WO 1999-US590
                           W
                                 19990112
     US 2000-610034
                           A2
                                 20000705
     US 2001-789017
                           A2
                                 20010220
     US 2001-288695P
                           Ρ
                                 20010503
     WO 2001-US32331
                           A1
                                 20011016
CLASS
 PATENT NO.
                 CLASS
                         PATENT FAMILY CLASSIFICATION CODES
 WO 9936086
                 ICM
                         A61K039-02
                         A61K039-385; C08B037-00
                 ICS
```

```
ECLA
                       A61K039/02; A61K039/385; C12P019/04
 WO 9936086
 US 2004126381
                 ECLA
                        A61K039/02; A61K039/102; A61K039/385; C12P019/04
                                                                              <--
                       A61K039/02; A61K039/385; C12P019/04
 US 2004115214
                 ECLA
     A conjugate vaccine for Moraxella catarrhalis comprising isolated
     lipooligosaccharide from which esterified fatty acids have been removed,
     to produce a detoxified lipooligosaccharide (dLOS), or from which lipid A
     has been removed, to produce a detoxified oligosaccharide (OS), which is
     linked to an immunogenic carrier. The immunogenic carrier is selected
     from the group consisting of UspA or CD derived from M. catarrhalis,
     tetanus toxoid, HMP derived from Haemophilus influenza, diphtheria toxoid,
     detoxified P. aeruginosa toxin A, cholera toxin, pertussis toxin,
     hepatitis B surface or core antigen, rotavirus VP 7 protein, CRM, CRM197,
     CRM3201 and respiratory syncytial virus F and G protein. The vaccine is
     useful for preventing otitis media and respiratory
     infections caused by M. catarrhalis in mammals, including humans.
ST
     Moraxella catarrhalis lipooligosaccharide vaccine conjugate; fatty acid
     lipid A removal vaccine
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (CD; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
     Proteins, specific or class
TT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(F; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (UspA; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
IT
     Glycoproteins, specific or class
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (VP7; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (adjuvant; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
IT.
     Immunostimulants
        (adjuvants; lipooligosaccharide-based vaccine for prevention
        of Moraxella (Branhamella) catarrhalis infections in mammals)
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (carrier; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
TT
        (chemical compds., linker; lipooligosaccharide-based vaccine for
        prevention of Moraxella (Branhamella) catarrhalis infections in
        mammals)
TΤ
     Toxins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (cholera; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
     Proteins, specific or class
IT
     RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); USES (Uses)
        (conjugates; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
ΤT
     Glycolipids
     RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); USES (Uses)
        (detoxified and conjugated; lipooligosaccharide-based vaccine for
        prevention of Moraxella (Branhamella) catarrhalis infections in
        mammals)
TT
     Toxins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (diphtheria, CRM and CRM197 and CRM3201; lipooligosaccharide-based
        vaccine for prevention of Moraxella (Branhamella) catarrhalis
        infections in mammals)
TТ
     Toxoids
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (diphtheria; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
IT
     Toxins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
```

(exotoxins; lipooligosaccharide-based vaccine for prevention of

```
Moraxella (Branhamella) catarrhalis infections in mammals)
IT
     Antigens
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hepatitis B core; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
TT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hepatitis B surface; lipooligosaccharide-based vaccine for prevention
        of Moraxella (Branhamella) catarrhalis infections in mammals)
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (high-mol.-weight; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
IT
     Carriers
        (immunogenic; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
IT
     Respiratory tract
        (infection; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
IT
     Drug delivery systems
        (injections, i.m.; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
IT
    Drug delivery systems
        (injections, s.c.; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
     Proteins, specific or class
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (linker; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
     Clostridium perfringens
IT
       Haemophilus influenzae
       Moraxella catarrhalis
     Pseudomonas aeruginosa
     Respiratory syncytial virus
     Rotavirus
       Vaccines
        (lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
     Fatty acids, processes
     Lipid A
     RL: REM (Removal or disposal); PROC (Process)
        (lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
TT
    G proteins (guanine nucleotide-binding proteins)
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
IT
     Lipid A
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (monophosphates, adjuvant; lipooligosaccharide-based vaccine for
        prevention of Moraxella (Branhamella) catarrhalis infections in
        mammals)
ΙT
    Drug delivery systems
        (mucosal; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
IT
    Drug delivery systems
        (nasal, intra-; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
TТ
    Ear
        (otitis, otitis media;
        lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
IT
    Drug delivery systems
        (parenterals; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pertussis; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
     Toxoids
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (tetanus; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (toxin A; lipooligosaccharide-based vaccine for prevention of Moraxella
```

Graser 10/030313 Page 50

```
(Branhamella) catarrhalis infections in mammals)
     Acids, biological studies
IT
     Bases, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (treatment; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
IT
     99-20-7D, Trehalose, dimycolate derivative
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (adjuvant; lipooligosaccharide-based vaccine for prevention of
        Moraxella (Branhamella) catarrhalis infections in mammals)
     60-32-2, .epsilon.-Aminohexanoic acid 1071-93-8, Adipic acid dihydrazide 24954-67-4, p-Nitrophenylethyl amine 32449-92-6, D-Glucuronolactone
TT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (linker; lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
     302-01-2, Hydrazine, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (lipooligosaccharide-based vaccine for prevention of Moraxella
        (Branhamella) catarrhalis infections in mammals)
             THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 7
RE
(1) Edebrink, P; CARBOHYDR RES 1995, V266(2), P237 HCAPLUS
(2) Gibson, B; WO 9853851 A 1998 HCAPLUS
(3) Gu, X; INFECTION AND IMMUNITY 1993, V61(5), P1873 HCAPLUS
(4) Gu, X; INFECTION AND IMMUNITY 1996, V64(10), P4047 HCAPLUS
(5) Gu, X; INFECTION AND IMMUNITY 1998, V66(5), P1891 HCAPLUS
(6) Kelly, J; ANALYTICAL BIOCHEMISTRY 1996, V233(1), P15 HCAPLUS
(7) Murphy, T; MICROBIOLOGICAL REVIEWS 1996, V60(2), P267 HCAPLUS
L42 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
AN
     1997:128048 HCAPLUS
DN
     126:211022
     Entered STN: 26 Feb 1997
ED
     Vaccines for nontypeable Haemophilus influenzae
TI
IN
    Green, Bruce A.; Zlotnick, Gary W.
     Praxis Biologics, Inc., USA
SO
    U.S., 24 pp., Cont.-in-part of U.S. Ser. No. 320, 971, abandoned.
    CODEN: USXXAM
рπ
     Patent
     English
LΑ
    ICM A61K039-102
IC
     ICS A61K039-385
NCL
     424256100
     15-2 (Immunochemistry)
CC
FAN.CNT 2
     PATENT NO.
                              DATE
                                            APPLICATION NO.
                         KIND
                                                                   DATE
     -----
                                -----
                         ----
                                            ------
                         Α
                                            US 1990-491466
    US 5601831
                               19970211
                                                                   19900309 <--
    CA 2047681
                        AA
                               19900910
                                            CA 1990-2047681
                                                                   19900309 <--
     CA 2047681
                         С
                                20000201
                       · A1
    EP 606921
                               19940720
                                            EP 1994-100492
                                                                   19900309 <--
                               20000802
    EP 606921
                         В1
        R: AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, LU, NL, SE
     ES 2063965 T3 19950116 ES 1990-905112
                                                                   19900309 <--
     AT 195076
                         E
                               20000815
                                            AT 1994-100492
                                                                   19900309 <--
                                            US 1995-447653
    US 5780601
                                                                   19950523 <--
                               19980714
                         А
     US 5955580
                         Α
                               19990921
                                            US 1995-449406
                                                                   19950523 <--
     US 6420134
                         В1
                                20020716
                                            US 1995-448097
                                                                   19950523 <--
PRAI US 1989-320971
                         В2
                                19890309
     EP 1990-905112
                         Α3
                                19900309 <--
    US 1990-491466
                         A3
                               19900309 <--
CLASS
                CLASS PATENT FAMILY CLASSIFICATION CODES
PATENT NO.
                _____
US 5601831
                ICM
                       A61K039-102
                 ICS
                       A61K039-385
                NCL
                        424256100
US 5601831
                ECLA
                       C07K014/285; C07K016/12A30; C07K019/00
EP 606921
                ECLA
                       C07K014/285
                                                                             <--
                        C07K014/285; C07K016/12A30; C07K019/00
US 5780601
                ECLA
US 5955580
                        C07K014/285; C07K016/12A30; C07K019/00
                                                                             <--
                 ECLA
                       C07K014/285; C07K016/12A30; C07K019/00
US 6420134
                ECLA
    Protein "e" of H. influenzae, a lipoprotein of approx. 28,000 daltons, has been purified and sequenced. Protein "e" and peptides or proteins having
```

```
a shared epitope, can be used to vaccinate against non-typable (and
typable) H. influenzae and to prevent otitis media
caused by H. influenzae. For this purpose, protein "e" or derivs. thereof
can be produced in native, synthetic or recombinant forms and can be
administered alone or in conjunction with other antigens of H. influenzae.
Protein "e" can also be used in multivalent vaccines designed for H.
influenzae and one or more other infectious organisms. Protein "e" was
isolated from Haemophilus cell envelopes and characterized, polyclonal
anti-protein "e" antiserum and monoclonal anti-protein "e" antibodies were
prepared, protein "e" gene was isolated and nucleotide sequence was determined
and mol. cloning of the gene was performed, bactericidal activity of
vaccine comprising protein "e" subunit was studied, and synergy of
anti-protein "e" with other antibodies were demonstrated.
vaccine Haemophilus influenzae protein e antibody
Proteins, specific or class
RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
(Preparation); USES (Uses)
   (E; protein "e" and gene of Haemophilus influenza and antibodies and
   vaccines for nontypeable Haemophilus influenzae)
Proteins, specific or class
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
   (OMP (outer membrane protein); protein "e" and gene of Haemophilus
   influenza and antibodies and vaccines for nontypeable Haemophilus
   influenzae)
Immunostimulants
   (adjuvants; protein "e" and gene of Haemophilus influenza and
   antibodies and vaccines for nontypeable Haemophilus influenzae)
Antibodies
RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
(Biological study); PREP (Preparation); USES (Uses)
   (bactericidal; protein "e" and gene of Haemophilus influenza and
   antibodies and vaccines for nontypeable Haemophilus influenzae)
Peptides, biological studies
Proteins, general, biological studies
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
   (carrier; protein "e" and gene of Haemophilus influenza and antibodies
   and vaccines for nontypeable Haemophilus influenzae)
Toxins
Toxoids
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (diphtheria; protein "e" and gene of Haemophilus influenza and
   antibodies and vaccines for nontypeable Haemophilus influenzae)
Lipoproteins
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
   (e; protein "e" and gene of Haemophilus influenza and antibodies and
   vaccines for nontypeable Haemophilus influenzae)
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
   (exotoxin A, Pseudomonas; protein "e" and gene of Haemophilus influenza
   and antibodies and vaccines for nontypeable Haemophilus influenzae)
Pseudomonas
   (exotoxin A; protein "e" and gene of Haemophilus influenza and
   antibodies and vaccines for nontypeable Haemophilus influenzae)
Gene microbial
RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
PRP (Properties); PUR (Purification or recovery); BIOL (Biological study);
PREP (Preparation)
   (for Haemophilus influenza protein "e"; protein "e" and gene of
   Haemophilus influenza and antibodies and vaccines for nontypeable
   Haemophilus influenzae)
Escherichia coli
   (heat labile toxin; protein "e" and gene of Haemophilus influenza and
   antibodies and vaccines for nontypeable Haemophilus influenzae)
Toxins
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
   (heat-labile, Escherichia coli; protein "e" and gene of Haemophilus influenza and antibodies and vaccines for nontypeable Haemophilus
   influenzae)
Antibodies
RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
(Biological study); PREP (Preparation); USES (Uses)
(monoclonal; protein "e" and gene of Haemophilus influenza and
   antibodies and vaccines for nontypeable Haemophilus influenzae)
   (otitis media; protein "e" and gene of Haemophilus
```

IT

IT

IT

TT

IT

IT

IT

TT

IT

IT

IT

IT

```
influenza and antibodies and vaccines for nontypeable Haemophilus
        influenzae)
TT
     Rotavirus
        (particles; protein "e" and gene of Haemophilus influenza and
        antibodies and vaccines for nontypeable Haemophilus influenzae)
TТ
     Bacterium (genus)
     DNA sequences
       Haemophilus influenzae
     Microorganism
       Moraxella catarrhalis
     Parasite
     Protein sequences
     Respiratory syncytial virus
     Staphylococcus aureus
     Streptococcus pneumoniae
     Streptococcus pyogenes
       Vaccines
     Virus
        (protein "e" and gene of Haemophilus influenza and antibodies and
        vaccines for nontypeable Haemophilus influenzae)
IT
     Opsonins
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (protein "e" and gene of Haemophilus influenza and antibodies and
        vaccines for nontypeable Haemophilus influenzae)
TT
     Antigens
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (protein "e" and gene of Haemophilus influenza and antibodies and
        vaccines for nontypeable Haemophilus influenzae)
     Oligosaccharides, biological studies
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (protein "e" and gene of Haemophilus influenza and antibodies and
        vaccines for nontypeable Haemophilus influenzae)
     Polysaccharides, biological studies
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (protein "e" and gene of Haemophilus influenza and antibodies and
        vaccines for nontypeable Haemophilus influenzae)
     Fusion proteins (chimeric proteins)
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (protein "e" and gene of Haemophilus influenza and antibodies and
        vaccines for nontypeable Haemophilus influenzae)
IT
     Toxins
     Toxoids
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (tetanus; protein "e" and gene of Haemophilus influenza and antibodies
        and vaccines for nontypeable Haemophilus influenzae)
     145110-32-3, Lipoprotein e (Haemophilus influenzae clone pPX513 gene hel
     precursor protein moiety reduced)
     RL: PRP (Properties)
        (amino acid sequence; protein "e" and gene of Haemophilus influenza and
        antibodies and vaccines for nontypeable Haemophilus influenzae)
     135622-17-2, DNA (Haemophilus influenzae type b strain Eagan clone pPX513
     lipoprotein e gene)
     RL: PRP (Properties)
        (nucleotide sequence; protein "e" and gene of Haemophilus influenza and
        antibodies and vaccines for nontypeable Haemophilus influenzae)
     82197-76-0, Polyribosylribitolphosphate
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (protein "e" and gene of Haemophilus influenza and antibodies and
        vaccines for nontypeable Haemophilus influenzae)
=> d all 130 tot
L30 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
AN
     2002:237317 HCAPLUS
DN
     136:261813
     Entered STN: 28 Mar 2002
ED
     Transferrin receptor-encoding genes from Haemophilus influenzae strains
     and their uses for diagnostics and medical treatment
    Loosmore, Sheena M.; Harkness, Robin E.; Schryvers, Anthony B.;
```

Graser 10/030313 Page 53

```
Chong, Pele; Gray-Owen, Scott; Yang, Yan-ping; Murdin, Andrew
     D.; Klein, Michel H.
     Aventis Pasteur Limited, Can.
PΑ
     U.S., 280 pp., Cont.-in-part of Ser. No. US 1995-483577, filed on 7 Jun
SO
     1995, now
     CODEN: USXXAM
DΤ
     Patent
T.A
     English
IC
     ICM A61K038-21
     ICS
         A61K038-16; A01N063-00; C12P019-34
NCL
    424256100
     15-2 (Immunochemistry)
CC
     Section cross-reference(s): 1, 3, 6
     PATENT NO.
                         KIND
                                DATE
                                            APPLICATION NO.
                                                                    DATE
                         ----
PΙ
     US 6361779
                          B1
                                20020326
                                            US 1996-649518
                                                                    19960517
     US 5922562
                          Α
                                19990713
                                             US 1994-337483
                                                                    19941108
                                             US 1995-483577
     US 6015688
                                20000118
                                                                    19950607
                          Α
                                             CA 1996-2223503
     CA 2223503
                          AA
                                19961219
                                                                    19960607
                                19961219
     WO 9640929
                          A2
                                             WO 1996-CA399
                                                                    19960607
                          А3
                                19970306
     WO 9640929
         W: AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE,
             ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS, LT,
             LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
             SG, SI
         RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR,
             IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN
                                             AU 1996-61177
                                                                    19960607
                                19961230
     AU 9661177
                          A1
     AU 716506
                          B2
                                20000224
     EP 833920
                          A2
                                19980408
                                             EP 1996-918543
                                                                    19960607
     EP 833920
                          Bl
                                20040818
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
     JP 11506335
                          Т2
                                19990608
                                             JP 1997-500057
                                                                    19960607
     JP 3516688
                          B2
                                20040405
     BR 9608482
                                20010731
                                             BR 1996-8482
                                                                    19960607
                          А
                                             AT 1996-918543
     AT 274059
                          E
                                20040915
                                                                    19960607
     US 2003088086
                          A1
                                20030508
                                             US 2002-43344
                                                                    20020114
PRAI US 1993-148968
                          B2
                                19931108
     US 1993-175116
                          B2
                                19931229
     US 1994-337483
                          A2
                                19941108
     US 1995-483577
                          A2
                                19950607
     US 1996-649518
                          Α
                                19960517
                                19960607
     WO 1996-CA399
CLASS.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
US 6361779
                 ICM
                        A61K038-21
                        A61K038-16; A01N063-00; C12P019-34
                 ICS
                        424256100
                 NCL
 US 6361779
                 ECLA
                        C07K014/285
 US 5922562
                 ECLA
                        C07K014/285
                 ECLA
                        C07K014/285
 US 6015688
                        C07K014/285
WO 9640929
                 ECT.A
US 2003088086
                ECLA
                        C07K014/285
    Purified and isolated genes are provided which encodes transferrin
     receptor proteins Tbp1 and/or Tbp2 of Haemophilus influenzae type b
     strains DL63, Eagan, MinnA, PAK12085, and SB33 and the non-typeable strains SB12, SB29, SB30, and SB32. The nucleic acid sequence may be used
     to produce peptides free of contaminants derived from bacteria normally
     containing the Tbp1 or Tbp2 proteins for purposes of diagnostics and medical
     treatment. Furthermore, the nucleic acid mol. may be used in the
     diagnosis of infection. Also provided are recombinant Tbpl or Tbp2 and
     methods for purification of the same. Live vectors expressing epitopes of
     transferrin receptor protein for vaccination are provided. Thus,
     poliovirus vectors incorporating the H. influenzae strain DL63 Tbp2 are
     neutralized by guinea-pig antisera raised against peptide LEGGFYGP,
     indicating that the viruses express this sequence in an antigenically
     recognizable form. Since H. influenzae Tbp2 is produced in low amts by
     Escherichia coli, the Eagan strain Tbp2 gene was truncated from its 3'-end
     using an Erase-a-base kit to produce a number of truncated analogs of Tbp2.
     The yield of Eagan rTbp2 is significantly increased by truncation of the
     C-terminal region of the protein. The infant rat model of bacteremia
     confirms the protective ability of anti-(truncated analogs of transferrin
     receptor protein Tbp2) antibodies even after removal of up to half of the
```

```
Tbp2 sequence.
ST
     transferrin receptor gene sequence Haemophilus; antigenicity transferrin
     receptor Haemophilus; vaccination transferrin receptor Haemophilus
IT
     Plasmid vectors
        (JD-1468-29 and JD-1424-2-8, for expression in Escherichia coli;
        transferrin receptor-encoding genes from Haemophilus influenzae strains
        and their uses for diagnostics and medical treatment)
TΤ
    Gene. microbial
      Transferrin receptors
     RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
    DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (Tbpl and Tbp2; transferrin receptor-encoding genes from Haemophilus
        influenzae strains and their uses for diagnostics and medical
        treatment)
IT
    Moraxella catarrhalis
        (antiserum cross-reactivity with; transferrin receptor-encoding genes
        from Haemophilus influenzae strains and their uses for diagnostics and
       medical treatment)
IT
    Immunoassay
        (enzyme, development and cross-reactivity of; transferrin
        receptor-encoding genes from Haemophilus influenzae strains and their
        uses for diagnostics and medical treatment)
IT
    Diagnosis
        (mol.; transferrin receptor-encoding genes from Haemophilus influenzae
        strains and their uses for diagnostics and medical treatment)
IT
    Escherichia coli
        (plasmid vectors JD-1468-29 and JD-1424-2-8 for expression in;
        transferrin receptor-encoding genes from Haemophilus influenzae strains
        and their uses for diagnostics and medical treatment)
IT
    Viral vectors
        (poliovirus type 1; transferrin receptor-encoding genes from
        Haemophilus influenzae strains and their uses for diagnostics and
        medical treatment)
IT
    DNA sequences
       Epitopes
      Haemophilus influenzae
    Molecular cloning
     Protein sequences
     Vaccines
        (transferrin receptor-encoding genes from Haemophilus influenzae
        strains and their uses for diagnostics and medical treatment)
     Promoter (genetic element)
     RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
     study); USES (Uses)
        (transferrin receptor-encoding genes from Haemophilus influenzae
        strains and their uses for diagnostics and medical treatment)
IT
    Human poliovirus 1
        (vector; transferrin receptor-encoding genes from Haemophilus
        influenzae strains and their uses for diagnostics and medical
        treatment)
     405178-16-7P
                    405178-17-8P
                                  405178-18-9P
                                                  405178-19-0P
                                                                 405178-20-3P
IT
     405178-21-4P
                   405178-23-6P
                                   405178-24-7P
                                                  405178-26-9P
                                                                 405178-28-1P
     405178-30-5P
                   405178-32-7P
    RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
    DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (amino acid sequence; transferrin receptor-encoding genes from
       Haemophilus influenzae strains and their uses for diagnostics and
       medical treatment)
    405178-36-1P
                   405178-37-2P
                                   405178-38-3P
                                                  405178-39-4P
                                                                405178-40-7P
IT
    405178-41-8P
                    405178-42-9P
                                   405178-43-0P
                                                  405178-44-1P
                                                                405178-45-2P
                   405178-47-4P
                                   405178-48-5P
    405178-46-3P
    RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); BIOL (Biological study); PREP (Preparation)
        (amino acid sequence; transferrin receptor-encoding genes from
       Haemophilus influenzae strains and their uses for diagnostics and
       medical treatment)
IT
    167769-62-2 167769-63-3
    RL: PAC (Pharmacological activity); PRP (Properties); BIOL (Biological
    study)
        (antigenic peptide epitope; transferrin receptor-encoding genes from
       Haemophilus influenzae strains and their uses for diagnostics and
       medical treatment)
    405178-12-3P
                   405178-13-4P
                                   405178-14-5P
                                                  405178-15-6P
                                                                 405178-22-5P
                   405178-27-0P
                                  405178-29-2P
    405178-25-8P
                                                  405178-31-6P
                                                                 405178-33-8P
```

```
405178-34-9P
                   405178-35-0P
     RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
     DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (nucleotide sequence; transferrin receptor-encoding genes from
        Haemophilus influenzae strains and their uses for diagnostics and
        medical treatment)
TT
     405180-53-2
                   405180-54-3
                                  405180-55-4
                                                 405180-56-5
                                                               405180-57-6
                                                 405180-68-9
     405180-58-7
                   405180-59-8
                                  405180-60-1
                                                               405180-69-0
     405180-70-3
                   405180-71-4
                                  405180-72-5
                                                 405180-73-6
                                                               405180-74-7
     405180-75-8
                   405180-76-9
                                  405180-77-0
                                                 405180-78-1
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; transferrin receptor-encoding genes
        from Haemophilus influenzae strains and their uses for diagnostics and
        medical treatment)
                   405180-07-6
TΤ
     405180-06-5
                                  405180-08-7
                                                 405180-09-8
                                                               405180-10-1
     405180-11-2
                   405180-12-3
                                  405180-13-4
                                                 405180-14-5
                                                               405180-15-6
     405180-16-7
                   405180-17-8
                                  405180-18-9
                                                 405180-19-0
                                                               405180-20-3
     405180-21-4
                   405180-22-5
                                  405180-23-6
                                                 405180-24-7
                                                               405180-25-8
     405180-26-9
                   405180-27-0
                                  405180-28-1
                                                 405180-29-2
                                                               405180-30-5
     405180-31-6
                   405180-32-7
                                  405180-33-8
                                                 405180-34-9
                                                               405180-35-0
                                  405180-38-3
     405180-36-1
                   405180-37-2
                                                 405180-39-4
                                                               405180-40-7
     405180-41-8
                   405180-42-9
                                  405180-43-0
                                                 405180-44-1
                                                               405180-45-2
     405180-46-3
                   405180-47-4
                                  405180-48-5
                                                 405180-49-6
                                                               405180-50-9
     405180-51-0
                   405180-52-1
                                  405180-61-2
                                                 405180-62-3
                                                               405180-63-4
     405180-64-5
                   405180-65-6
                                  405180-66-7
                                                 405180-67-8
     RL: PRP (Properties)
        (unclaimed protein sequence; transferrin receptor-encoding genes from
        Haemophilus influenzae strains and their uses for diagnostics and
        medical treatment)
IT
     161228-75-7
                   229032-30-8
                                  229032-31-9
                                                 229032-32-0
                                                               229032-33-1
     229032-34-2
                   229032-35-3
                                  229032-36-4
                                                 229032-37-5
                                                               229032-38-6
     229032-39-7
                   229032-40-0
                                  229032-41-1
                                                 229032-42-2
                                                               229032-43-3
     229032-44-4
                   229032-45-5
                                  229032-46-6
                                                 229032-47-7
                                                               229032-48-8
     229157-61-3
                   229157-62-4
                                  229157-63-5
                                                 229157-64-6
                                                               229157-65-7
     404572-60-7
                   404572-61-8
                                  404572-62-9
                                                 404572-63-0
                                                               404572-64-1
     404572-65-2
                  404572-66-3
                                  404572-67-4
                                                 404572-68-5
                                                               404572-69-6
     404572-70-9
                  404572-72-1
                                  404572-74-3
                                                 404572-76-5
                                                               404572-78-7
     404572-80-1
                  404572-82-3
                                  404572-84-5
     RL: PRP (Properties)
        (unclaimed sequence; transferrin receptor-encoding genes from
        Haemophilus influenzae strains and their uses for diagnostics and
        medical treatment)
RE.CNT
              THERE ARE 39 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
(1) Anon; WO 9412641 1994 HCAPLUS
(2) Anon; WO 9513370 1995 HCAPLUS
(3) Barber; US 5194254 A 1993 HCAPLUS
(4) Frasch; US 4601903 A 1986
(5) Ghrayeb; Embo J 1984, V3, P2437 HCAPLUS
(6) Gordon; US 4496538 A 1985 HCAPLUS
(7) Gray-Owen; Infect Immun 1995, V4, P1201
(8) Griffiths; Fems Microbiol Lett 1993, V109(1), P85 HCAPLUS
(9) Holland; Infect Immun 1992, V60, P2986 HCAPLUS
(10) Hopp, T; Journal of Immunological Methods 1986, V88, P1 HCAPLUS
(11) Itakura; Science 1977, V198, P1056 HCAPLUS
(12) Jarosik; Infection and Immunity 1994, V62, P2470 HCAPLUS
(13) Legrain; Gene 1993, V130, P73 HCAPLUS
(14) Lockhoff; Chem Ind Ed Engl 1991, V30, P1611 (15) Lockoff; US 4855283 A 1989 HCAPLUS
(16) Mickelsen; Infect Immun 1981, V33, P555 HCAPLUS
(17) Milich; US 4599230 A 1986 HCAPLUS
(18) Milich; US 4599231 A 1986 HCAPLUS
(19) Moloney; US 4258029 A 1981 HCAPLUS
(20) Morton; Infection and Immunity 1993, V61, P4033 HCAPLUS
(21) Murdin; J Gen Viral 1992, V73, P607 HCAPLUS
(22) Murdin; Microbial Pathogenesis 1991, V10, P27 HCAPLUS
(23) Nixon-George; J Immunol 1990, V14, P4798
(24) Ogunnariwo; Avian Dis 1992, V36, P655 MEDLINE (25) O'Hagan; Clin Pharmokinet 1992, V22, P1 MEDLINE
(26) Panezutti; Infection and Immunity 1993, V61, P1867 HCAPLUS
(27) Roosi-Campos; Vaccine 1992, V10, P512
(28) Sambrook; Molecular Cloning, A Lab Manual V3, P16.2
(29) Schryvers; US 5141743 A 1992 HCAPLUS
(30) Schryvers; J Infect Dis 1992, V165(suppl 1), PS103
(31) Schryvers; Molec Microbiol 1988, V2, P467 HCAPLUS
```

```
(32) Schyrvers; Can J Microbiol 1989, V35, P409
(33) Schyvers; Med Microbiol 1989, V29, P121
(34) Short; Nucl Acids Res 1988, V16, P7583 HCAPLUS
(35) Studier; US 4952496 A 1990 HCAPLUS
(36) Ulmer; Curr Opinion Invest Drugs 1993, V2(9), P983
(37) Vanderwerf; Proc Natl Acad Sci 1986, V83, P2330 HCAPLUS
(38) Vyas; US 4596792 A 1986 HCAPLUS
(39) Wilton; FEMS Microbiology Letters 1993, V107, P59 HCAPLUS
L30 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
     2002:213747 HCAPLUS
AN
DN
     136:242991
ED
     Entered STN: 21 Mar 2002
     Transferrin receptor-encoding genes from Haemophilus influenzae strains
TI
     and their uses for diagnostics and medical treatment
IN
     Loosmore, Sheena M.; Harkness, Robin E.; Schryvers, Anthony B.;
     Chong, Pele; Gray-Owen, Scott; Yang, Yan-Ping; Murdin, Andrew
     D.; Klein, Michel H.
     Aventis Pasteur Limited, Can.
PΑ
     U.S., 264 pp., Cont.-in-part of U.S. Ser. No. 175,116, abandoned.
SO
     CODEN: USXXAM
DΤ
     Patent
     English
LA
     ICM C12N001-21
TC
     ICS C12N005-10; C12N015-03; C12N015-63
     435252300
NCL
CC
     3-3 (Biochemical Genetics)
     Section cross-reference(s): 1, 6, 15
FAN.CNT 5
     PATENT NO.
                         KIND DATE
                                             APPLICATION NO.
                                                                     DATE
     US 6358727
                                20020319
                         B1
                                             US 1996-637654
                                                                    19960805
PΙ
                                19950518
                                             WO 1994-CA616
     WO 9513370
                          Al
                                                                    19941107
         W: AU, BR, CA, CN, FI, JP, KR, NO, NZ, RU, UA, US
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
                                19931108
PRAI US 1993-148968
                          B2
     US 1993-175116
                          B2
                                19931229
     WO 1994-CA616
                                19941107
                          W
CLASS
                 CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
 US 6358727
                 ICM
                        C12N001-21
                        C12N005-10; C12N015-03; C12N015-63
                 ICS
                        435252300
                 NCL
US 6358727
                        C07K014/285
                 ECLA
WO 9513370
                 ECLA
                       C07K014/285
     Purified and isolated genes are provided which encodes transferrin
     receptor proteins Tbpl and/or Tbp2 of Haemophilus influenzae type b
     strains DL63, Eagan, MinnA, PAK12085, and SB33 and the non-typeable strains SB12, SB29, SB30, and SB32. The nucleic acid sequence may be used
     to produce peptides free of contaminants derived from bacteria normally
     containing the Tbp1 or Tbp2 proteins for purposes of diagnostics and medical
     treatment. Furthermore, the nucleic acid mol. may be used in the
     diagnosis of infection. Also provided are recombinant Tbp1 or Tbp2 and
     methods for purification of the same. Live vectors expressing epitopes of
     transferrin receptor protein for vaccination are provided. Thus,
     poliovirus vectors incorporating the H. influenzae strain DL63 Tbp2 are
     neutralized by guinea-pig antisera raised against peptide LEGGFYGP,
     indicating that the viruses express this sequence in an antigenically
     recognizable form.
     transferrin receptor gene sequence Haemophilus; antigenicity transferrin
     receptor Haemophilus; vaccination transferrin receptor Haemophilus
TT
     Plasmid vectors
        (JD-1468-29 and JD-1424-2-8, for expression in Escherichia coli;
        transferrin receptor-encoding genes from Haemophilus influenzae strains
        and their uses for diagnostics and medical treatment)
IT
     Gene, microbial
       Transferrin receptors
     RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
     DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (Tbp1 and Tbp2; transferrin receptor-encoding genes from Haemophilus
        influenzae strains and their uses for diagnostics and medical
        treatment)
IT
    Moraxella catarrhalis
        (antiserum cross-reactivity with; transferrin receptor-encoding genes
```

```
from Haemophilus influenzae strains and their uses for diagnostics and
       medical treatment)
TΤ
    Immunoassay
        (enzyme, development and cross-reactivity of; transferrin
        receptor-encoding genes from Haemophilus influenzae strains and their
       uses for diagnostics and medical treatment)
    Diagnosis
IT
        (mol.; transferrin receptor-encoding genes from Haemophilus influenzae
        strains and their uses for diagnostics and medical treatment)
     Escherichia coli
IT
        (plasmid vectors JD-1468-29 and JD-1424-2-8 for expression in;
        transferrin receptor-encoding genes from Haemophilus influenzae strains
        and their uses for diagnostics and medical treatment)
TТ
    Viral vectors
        (poliovirus type 1; transferrin receptor-encoding genes from
       Haemophilus influenzae strains and their uses for diagnostics and
        medical treatment)
IT
    DNA sequences
      Epitopes
      Haemophilus influenzae
    Molecular cloning
     Protein sequences
     Vaccines
        (transferrin receptor-encoding genes from Haemophilus influenzae
        strains and their uses for diagnostics and medical treatment)
     Promoter (genetic element)
     RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
     study); USES (Uses)
        (transferrin receptor-encoding genes from Haemophilus influenzae
        strains and their uses for diagnostics and medical treatment)
IT
    Human poliovirus 1
        (vector; transferrin receptor-encoding genes from Haemophilus
        influenzae strains and their uses for diagnostics and medical
        treatment)
    404796-06-1P
                   404796-07-2P
                                   404796-08-3P
                                                  404796-09-4P
                   404796-12-9P
                                   404796-13-0P
                                                 404796-14-1P
     404796-11-8P
                                                                404796-15-2P
     404796-16-3P
                   404796-17-4P
    RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
     DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (amino acid sequence; transferrin receptor-encoding genes from
       Haemophilus influenzae strains and their uses for diagnostics and
       medical treatment)
IT
    167769-62-2 167769-63-3
    RL: PAC (Pharmacological activity); PRP (Properties); BIOL (Biological
        (antigenic peptide epitope; transferrin receptor-encoding genes from
       Haemophilus influenzae strains and their uses for diagnostics and
       medical treatment)
                                   404795-96-6P
                                                  404795-97-7P
TT
    404795-94-4P
                   404795-95-5P
                                                                 404795-98-8P
     404795-99-9P
                    404796-00-5P
                                   404796-01-6P
                                                 404796-02-7P
                                                                404796-03-8P
    404796-04-9P
                   404796-05-0P
    RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);
    DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (nucleotide sequence; transferrin receptor-encoding genes from
       Haemophilus influenzae strains and their uses for diagnostics and
       medical treatment)
    404818-86-6
                  404818-87-7
                                 404818-88-8
TT
                                               404818-89-9
                                                             404818-90-2
     404818-91-3
                   404818-92-4
                                 404818-93-5
                                               404819-01-8
                                                             404819-02-9
                                               404819-06-3
    404819-03-0
                  404819-04-1
                                 404819-05-2
                                                             404819-07-4
                                 404819-10-9
                                               404819-11-0
    404819-08-5
                  404819-09-6
    RL: PRP (Properties)
        (unclaimed nucleotide sequence; transferrin receptor-encoding genes
        from Haemophilus influenzae strains and their uses for diagnostics and
       medical treatment)
IT
    404818-39-9
                  404818-40-2
                                 404818-41-3
                                                             404818-43-5
                                               404818-42-4
    404818-44-6
                  404818-45-7
                                 404818-46-8
                                               404818-47-9
                                                             404818-48-0
                  404818-50-4
                                 404818-51-5
    404818-49-1
                                               404818-52-6
                                                             404818-53-7
                  404818-55-9
                                 404818-56-0
                                               404818-57-1
                                                             404818-58-2
    404818-54-8
    404818-59-3
                  404818-60-6
                                 404818-61-7
                                               404818-62-8
                                                             404818-63-9
    404818-64-0
                  404818-65-1
                                 404818-66-2
                                               404818-67-3
                                                             404818-68-4
                  404818-70-8
                                 404818-71-9
                                               404818-72-0
    404818-69-5
                                                             404818-73-1
                  404818-75-3
    404818-74-2
                                 404818-76-4
                                               404818-77-5
                                                             404818-78-6
    404818-79-7
                  404818-80-0
                                 404818-81-1
                                               404818-82-2
                                                             404818-83-3
    404818-84-4
                  404818-85-5
                                 404818-94-6
                                               404818-95-7
                                                             404818-96-8
```

Graser 10/030313 Page 58

404818-97-9 404818-98-0 404818-99-1 404819-00-7 RL: PRP (Properties) (unclaimed protein sequence; transferrin receptor-encoding genes from Haemophilus influenzae strains and their uses for diagnostics and medical treatment) TT 161228-75-7 229032-30-8 229032-31-9 229032-32-0 229032-33-1 229032-34-2 229032-35-3 229032-36-4 229032-37-5 229032-38-6 229032-39-7 229032-40-0 229032-41-1 229032-42-2 229032-43-3 229032-44-4 229032-45-5 229032-46-6 229032-47-7 229032-48-8 229157-61-3 229157-62-4 229157-63-5 229157-64-6 229157-65-7 404572-61-8 404572-62-9 404572-63-0 404572-60-7 404572-64-1 404572-66-3 404572-72-1 404572-65-2 404572-67-4 404572-68-5 404572-69-6 404572-70-9 404572-74-3 404572-76-5 404572-78-7 404572-82-3 404572-80-1 404572-84-5 RL: PRP (Properties) (unclaimed sequence; transferrin receptor-encoding genes from Haemophilus influenzae strains and their uses for diagnostics and medical treatment) THERE ARE 60 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT RE (1) Anon; WO 9306861 1993 HCAPLUS (2) Anon; WO 9308283 1993 HCAPLUS (3) Barcak; Methods Enzymol 1991, V204, P321 HCAPLUS (4) Berber; US 5194254 A 1993 HCAPLUS (5) Berkowitz; J Pediatr 1987, V110, P509 MEDLINE (6) Black; Pediatr Infect Dia J 1991, V10, P97 MEDLINE (7) Bluestone, N; Engl J Med 1982, V306, P1399 (8) Cheng; Nature 1978, V375, P615 (9) Chou; Annual Reviews of Biochemistry 1978, V47, P251 HCAPLUS (10) Claesson; J Pediatr 1989, V114, P97 MEDLINE (11) Cornelissen; J of Bacteriology 1992, V174, P5788 HCAPLUS (12) Danve; Vaccine 1993, V11, P1214 HCAPLUS (13) Derea; Nature 1989, V342, P651 (14) Freach; US 4601903 A 1986 (15) George; Macromolecular Sequencing and Synthesis 1988, P127 (16) Gerlach; Infection and Immunity 1992, V60, P3253 HCAPLUS (17) Ghrayeb; The Embo J 1984, V3, P2437 HCAPLUS (18) Goeddel; Nature 1979, V281, P544 HCAPLUS (19) Goldberg; Mol Microbiology 1992, V6, P2407 HCAPLUS (20) Gordon: US 4496538 A 1985 HCAPLUS (21) Gray-Owen; Infect Immun 1995, V63(4), P1201 HCAPLUS (22) Gray-Owen; Microbial Pathogenesis 1993, V14 MEDLINE (23) Griffiths; Fems Microbiol Lett 1993, V109(1), P85 HCAPLUS (24) Harkness; J Bacteriol 1992, V174, P2425 HCAPLUS (25) Holland; FEMS, Microbiology Letters 1991, V77, P283 HCAPLUS (26) Holland; Infection and Immunity 1992, V60, P2986 HCAPLUS (27) Hopp, T; Journal of Immunological Methods 1986, V88, P1 HCAPLUS (28) Itakura; Science 1977, V198, P1058 (29) Jaroaik; Infection and Immunity 1994, V62, P2470 (30) Legrain; Gene 1993, V130, P73 HCAPLUS (31) Lockhoff; US 4855283 A 1989 HCAPLUS (32) Lockhoff; Chem Int Ed Engl 1991, V30, P1611 (33) McGeoch; J Gen Virol 1988, V69, P1531 HCAPLUS (34) Mickelsen; Infect Immun 1981, V33, P555 HCAPLUS (35) Millich; US 4599230 A 1986 HCAPLUS (36) Millich; US 4599231 A 1986 HCAPLUS (37) Moloney; US 4258029 A 1981 HCAPLUS (38) Morton; Infection and Immunity 1993, V61, P4033 HCAPLUS (39) Murdin; J Gen Viral 1992, V73, P607 HCAPLUS (40) Murdin; Microbial Pathogenesis 1991, V10, P27 HCAPLUS (41) Nixon-George; J Immunol 1990, V14, P4798 (42) Ogunnariwo; Avian Dis 1992, V38, P855 (43) O'Hagen; Clin Pharmokinet 1992, V22, P1 (44) Panazutti; Infection and Immunity 1993, V61, P1867 (45) Poulsen; Molecular Microbiology 1992, V6, P895 HCAPLUS (46) Roosi-Campos; Vaccine 1992, V10, P512 (47) Sambrook; Molecular Biology, A Laboratory Manual 2nd Ed 1989, V3, P16.2 (48) Schryvers; US 5141743 A 1992 HCAPLUS (49) Schryvers; Can J Microbiol 1989, V35, P409 HCAPLUS (50) Schryvers; J Infect Dis 1992, V165(suppl 1), PS103 (51) Schryvers; Molec Microbiol 1988, V2, P467 HCAPLUS (52) Schyvers; Med Microbiol 1989, V29, P121 (53) Short; Nucl Acids Res 1988, V16, P7533 (54) Stevenson; Infection and Immunity 1992, V60(8), P2391 (55) Studier; US 4952498 A 1990 (56) Thomas; Methods in Enzymology 1990, V182, P499 HCAPLUS

```
(57) Ulmer; Curr Opinion Invest Drugs 1993, V2(9), P983
(58) Vandarwerf; Proc Natl Acad Sci 1986, V83, P2330
(59) Vyae; US 4596792 A 1986 HCAPLUS
(60) Weismuller; Vaccine 1989, V8, P29
1,30
    ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
     2001:63846 HCAPLUS
AN
DN
     134:120915
     Entered STN: 26 Jan 2001
ΤI
     Multicomponent vaccine to protect against disease caused by Haemophilus
     influenzae and Moraxella catarrhalis
     Loosmore, Sheena M.; Yang, Yan-Ping; Klein, Michel H.;
IN
     Sasaki, Ken
PA
     Connaught Laboratories Limited, Can.
so
     PCT Int. Appl., 58 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
     ICM A61K039-00
IC
     63-3 (Pharmaceuticals)
CC
     Section cross-reference(s): 15
FAN.CNT 1
                          KIND
                                 DATE
                                              APPLICATION NO.
                                                                      DATE
     PATENT NO.
                          ----
ΡI
     WO 2001005424
                           A2
                                 20010125
                                              WO 2000-CA811
                                                                      20000711
     WO 2001005424
                                 20010802
                           АЗ
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
             HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
             LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
             SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
             YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     US 6391313
                           B1
                                 20020521
                                              US 1999-353617
                                                                      19990715
     CA 2378862
                                              CA 2000-2378862
                                                                      20000711
                           AΑ
                                 20010125
     EP 1200122
                           A2
                                 20020502
                                              EP 2000-945494
                                                                      20000711
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL
                                              AU 2000-59586
     AU 767096
                                 20031030
                                                                      20000711
                           B2
PRAI US 1999-353617
                           Α
                                 19990715
    WO 2000-CA811
                                 20000711
CLASS
PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
                 ----
WO 2001005424
                 ICM
                         A61K039-00
US 6391313
                 ECLA
                        A61K039/116
    A multi-valent immunogenic composition confers protection on an immunized host
     against infection caused by both Haemophilus influenzae and Moraxella
     catarrhalis. Such composition comprises at least four antigens comprising at
     least one antigen from Haemophilus influenzae, and at least one antigen
     from Moraxella catarrhalis. Three of the antigens are adhesins. High
     mol. weight (HMW) proteins and Haemophilus influenzae adhesin (Hia) proteins
     of non-typeable Haemophilus and a 200 kDa outer membrane protein of
     Moraxella catarrhalis comprise the adhesin components while the other
    antigen is a non-proteolytic analog of Hin47 protein. Each component does not impair the immunogenicity of the others. The multi-valent immunogenic
     composition may be combined with DTP component vaccines, which may also include
     non-virulent poliovirus and PRP-T, to provide a component vaccine without
     impairment of the immunogenic properties of the other antigens.
ST
     adhesin antigen vaccine Haemophilus Moraxella
     Proteins, specific or class
IT
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
        (HMW1; multicomponent vaccine to protect against disease caused by
        Haemophilus influenzae and Moraxella catarrhalis)
     Proteins, specific or class
    RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
        (HMW2; multicomponent vaccine to protect against disease caused by
        Haemophilus influenzae and Moraxella catarrhalis)
```

```
Proteins, specific or class
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
        (Hin47; multicomponent vaccine to protect against disease caused by
        Haemophilus influenzae and Moraxella catarrhalis)
     Proteins, specific or class
TT
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
        (Hsf (Haemophilus surface fibril); multicomponent vaccine to protect
        against disease caused by Haemophilus influenzae and Moraxella
        catarrhalis)
     Proteins, specific or class
IT
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
        (OMP (outer membrane protein); multicomponent vaccine to protect
        against disease caused by Haemophilus influenzae and Moraxella
        catarrhalis)
TT
     Immunostimulants
        (adjuvants; multicomponent vaccine to protect against disease caused by
        Haemophilus influenzae and Moraxella catarrhalis)
     Agglutinins and Lectins
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU
     (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
        (agglutinogens; multicomponent vaccine to protect against disease
        caused by Haemophilus influenzae and Moraxella catarrhalis)
     Adhesins
IT
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
        (antigenic; multicomponent vaccine to protect against disease caused by
        Haemophilus influenzae and Moraxella catarrhalis)
     Toxoids
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU
     (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
        (diphtheria; multicomponent vaccine to protect against disease caused by Haemophilus influenzae and Moraxella catarrhalis)
    Organelle
TΤ
        (fibril, surface; multicomponent vaccine to protect against disease
        caused by Haemophilus influenzae and Moraxella catarrhalis)
IT
     Hemagglutinins
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
        (filamentous; multicomponent vaccine to protect against disease caused
        by Haemophilus influenzae and Moraxella catarrhalis)
TT
     Chinchilla
       Haemophilus influenzae
     Molecular cloning
     Molecular weight distribution
       Moraxella catarrhalis
     Polyacrylamide gel electrophoresis
     Vaccines
        (multicomponent vaccine to protect against disease caused by
        Haemophilus influenzae and Moraxella catarrhalis)
IT
     Antigens
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PROC (Process); USES (Uses)
        (multicomponent vaccine to protect against disease caused by
        Haemophilus influenzae and Moraxella catarrhalis)
    Heat-shock proteins
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); THU (Therapeutic use); BIOL (Biological
     study); OCCU (Occurrence); PROC (Process); USES (Uses)
```

(non-proteolytic; multicomponent vaccine to protect against disease

```
caused by Haemophilus influenzae and Moraxella catarrhalis)
IT
     Human poliovirus
         (non-virulent; multicomponent vaccine to protect against disease caused
         by Haemophilus influenzae and Moraxella catarrhalis)
IT
         (otitis, otitis media; multicomponent vaccine to protect against
         disease caused by Haemophilus influenzae and Moraxella catarrhalis)
     Agglutinins and Lectins
TT
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (pertactins; multicomponent vaccine to protect against disease caused
        by Haemophilus influenzae and Moraxella catarrhalis)
     Toxoids
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU
      (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
        (pertussis; multicomponent vaccine to protect against disease caused by Haemophilus influenzae and Moraxella catarrhalis)
TT
     Mutation
         (substitution; multicomponent vaccine to protect against disease caused
         by Haemophilus influenzae and Moraxella catarrhalis)
IT
     Toxoids
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (tetanus; multicomponent vaccine to protect against disease caused by
         Haemophilus influenzae and Moraxella catarrhalis)
     9001-92-7, Proteinase
     RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BSU (Biological study, unclassified); BIOL (Biological
     study); OCCU (Occurrence)
         (activity levels; multicomponent vaccine to protect against disease
         caused by Haemophilus influenzae and Moraxella catarrhalis)
     7784-30-7, Aluminum phosphate
                                       21645-51-2, Aluminum hydroxide, biological
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
         (adjuvant; multicomponent vaccine to protect against disease caused by
        Haemophilus influenzae and Moraxella catarrhalis)
IT
     151-21-3, Sds, analysis
     RL: ARU (Analytical role, unclassified); ANST (Analytical study)
         (multicomponent vaccine to protect against disease caused by
        Haemophilus influenzae and Moraxella catarrhalis)
=> b wpix
FILE 'WPIX' ENTERED AT 13:10:45 ON 16 DEC 2004
COPYRIGHT (C) 2004 THE THOMSON CORPORATION
                              13 DEC 2004
FILE LAST UPDATED:
                                                  <20041213/UP>
MOST RECENT DERWENT UPDATE:
                                  200480
                                                   <200480/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE
>>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE,
    PLEASE VISIT:
 http://www.stn-international.de/training center/patents/stn guide.pdf <<<
>>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE
    http://thomsonderwent.com/coverage/latestupdates/
>>> FOR INFORMATION ON ALL DERWENT WORLD PATENTS INDEX USER .
    GUIDES, PLEASE VISIT:
    http://thomsonderwent.com/support/userguides/
>>> NEW! FAST-ALERTING ACCESS TO NEWLY-PUBLISHED PATENT
    DOCUMENTATION NOW AVAILABLE IN DERWENT WORLD PATENTS INDEX
    FIRST VIEW - FILE WPIFV.
    FOR FURTHER DETAILS: http://www.thomsonderwent.com/dwpifv <<<
>>> NEW DISPLAY FORMAT HITSTR ADDED ALLOWING DISPLAY OF
    HIT STRUCTURES WITHIN THE BIBLIOGRAPHIC DOCUMENT <><
```

>>> SMILES and ISOSMILES strings are no longer available as Derwent Chemistry Resource display fields <<<

=> d all 154 tot L54 ANSWER 1 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN 2004-774372 [76] WPIX AN DNC C2004-271112 New immunogenic composition comprises a major outer membrane protein of a strain of Chlamydia pneumoniae, and a 76 kDa protein of a strain of C. pneumoniae, useful as a vaccine for treating or preventing Chlamydia infections. DC B04 D16 DUNN, P L; MURDIN, A D ΤN PA (AVET) AVENTIS PASTEUR LTD CYC US 6811783 B1 20041102 (200476)\* 31 A61K039-02 PΙ ADT US 6811783 B1 US 1999-391606 19990907 PRAI US 1999-391606 19990907 ICM A61K039-02 ICS A61K039-00; C07H021-04; C07K001-00 AB 6811783 B UPAB: 20041125 US NOVELTY - An immunogenic composition comprises a first plasmid vector comprising a first nucleotide sequence encoding a major outer membrane protein (MOMP) of a strain of Chlamydia pneumoniae, and a second plasmid vector comprising a second nucleotide sequence encoding a 76 kDa protein of a strain of C. pneumoniae, is new. DETAILED DESCRIPTION - An immunogenic composition comprises a first plasmid vector comprising a first nucleotide sequence encoding a major outer membrane protein of a strain of C. pneumoniae, the first nucleotide sequence is selected from 3 sequences comprising 1426, 1301, or 1101 bp (SEQ ID NO. 12, 13, or 14) or encoding a MOMP having an amino acid sequence comprising 394 or 367 amino acids (SEQ ID NO. 15 or 16), and a first promoter sequence operatively coupled to the first nucleotide sequence for expression of the MOMP in a host; and a second plasmid vector comprising a second nucleotide sequence encoding a 76 kDa protein of a strain of C. pneumoniae, the second nucleotide sequence is selected from 4 sequences comprising 2545, 651, 1470, or 1389 bp (SEQ ID NO. 1, 2, 3, or 4), and a second promoter sequence operatively coupled to the second nucleotide sequence for expression of the 76 kDa protein in a host; and a pharmaceutical carrier. All sequences are defined in the specification. ACTIVITY - Antibacterial. MECHANISM OF ACTION - Vaccine. Groups of 7-9 week old male Balb/c mice were immunized intramuscularly (i.m.) and intranasally (i.n.) with plasmids pCA76kDa and pCAMOMP containing the coding sequences of C. pneumoniae 76 kDa and MOMP, respectively. Saline or plasmid vectors containing non-protective inserted chlamydial genes were given to groups of control animals. Results showed an increased protection afforded by the combination of the two constructs. It showed that mice immunized intramuscularly and intranasally with both pCA76kDa and pCAMOMP had chlamydial lung titers less than 6700 in 6 of 6 cases, where the range of values for control mice with saline were 15000-106100 IFU/lung in 20 out of 23 cases, and 12600-80600 IFU/lung in 11 out of 12 cases for mice immunized with the vectors containing non-protective genes. USE - The immunogenic composition is useful as a vaccine for immunizing a host against disease caused by infection with a strain of Chlamydia. It is also useful for treating or preventing Chlamydia infections. Dwg.0/5 FS CPI AB; DCN FA MC CPI: B04-E08; B14-A01A; B14-S03; B14-S09; B14-S11B; D05-H07 ANSWER 2 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN L54 AN 2001-648559 [74] WPIX DNN N2001-484575 DNC C2001-191446 TI Novel polypeptides from Chlamydia pneumoniae and genes encoding the polypeptide, useful for immunization of host e.g. human against disease caused by infection by a strain of Chlamydia.

DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J
(AVET) AVENTIS PASTEUR LTD; (DUNN-I) DUNN P; (MURD-I) MURDIN A

D; (OOME-I) OOMEN R P; (WANG-I) WANG J 95 WO 2001075114 A2 20011011 (200174)\* EN

DC

IN PA

ΡI

B04 D16 S03

Search done by Noble Jarrell

C12N015-31

90

```
Graser 10/030313
       RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TR TZ UG ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK
            DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ
            LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD
            SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
                     A 20011015 (200209)
                                                       C12N015-31
    AU 2001048178
    US 2002082402
                    A1 20020627 (200245)
                                                       C07H021-02
                    A1 20031204 (200380)
    US 2003224004
                                                       A61K039-00
    WO 2001075114 A2 WO 2001-CA462 20010404; AU 2001048178 A AU 2001-48178
     20010404; US 2002082402 Al Provisional US 2000-194477P 20000404, US
     2001-824588 20010403; US 2003224004 Al Provisional US 2000-194477P
     20000404, Cont of US 2001-824588 20010403, US 2003-359289 20030206
FDT AU 2001048178 A Based on WO 2001075114
PRAI US 2000-194477P
                          20000404; US 2001-824588
                                                          20010403:
    US 2003-359289
                          20030206
IC
    ICM A61K039-00; C07H021-02; C12N015-31
         A61K039-118; A61K039-38; A61K039-40;
          C07H021-04; C07K001-00; C07K014-00; C07K014-295; C07K016-12;
          C07K017-00; C12N015-11; C12N015-62; C12Q001-68; G01N033-53;
          G01N033-68
    WO 200175114 A UPAB: 20011217
     NOVELTY - A transmembrane polypeptide from Chlamydia, preferably C.
    pneumoniae comprising a 579 residue amino acid sequence, fully defined in
     the specification, an immunogenic fragment of at least 12 consecutive
     amino acids of S1, or a polypeptide modified without loss of
     immunogenicity and having at least 75 % identity to them, is new.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the
     following:
          (1) a nucleic acid molecule (II) comprising a sequence encoding (I),
     a 1940 nucleotide sequence (S2), fully defined in the specification, a
     sequence encoding a polypeptide encoded by S2, a sequence comprising at
    least 38 consecutive nucleotides of them, or a sequence encoding a
    polypeptide having at least 75 % identity to a polypeptide encoded by S2;
          (2) a nucleic acid molecule (IIa) comprising a sequence which is
     antisense to (II);
          (3) a nucleic acid molecule (IIb) comprising a sequence encoding a
     fusion protein (FP) comprising a polypeptide encoded by (II) and a second
    polypeptide;
          (4) a vaccine (IIIa) comprising a vaccine vector and at least one
     first nucleic acid encoding (I) or FP, which is capable of being
    expressed, and optionally the vaccine comprises a second nucleic acid
     encoding and capable of expressing an additional polypeptide which
    enhances the immune response to the polypeptide expressed by the first
    nucleic acid;
          (5) a vaccine (IIIb) comprising (II)-(IIb) and a vaccine vector,
    where (II) - (IIb) is expressed as a polypeptide, and optionally the vaccine
     comprises a second nucleic acid encoding an additional polypeptide which
    enhances the immune response to the polypeptide expressed by (II)-(IIb); (6) a pharmaceutical composition (PC) comprising (II)-(IIb), (IIIa)
    or (IIIb):
          (7) a unicellular host (IV) transformed with (II)-(IIb);
          (8) an isolated nucleic acid probe of 5-100 nucleotides which
    hybridizes under stringent conditions to S2, or its complement or
    antisense sequence;
          (9) an isolated primer of 10-40 nucleotides which hybridizes under
    stringent conditions to S2, or its complement or antisense sequence;
          (10) a polypeptide (Ia) encoded by (II)-(IIb);
          (11) a fusion protein (FP) comprising (I) or (Ia), and a second
    polypeptide;
          (12) producing (I) and FP;
          (13) an antibody (Ab) against (I) or FP;
(14) a vaccine (IIIc) comprising (I), a polypeptide encoded by (II),
    or FP comprising (I) and a second polypeptide, and optionally comprising
    an additional polypeptide which enhances the immune response to the first
    polypeptide;
          (15) a vaccine (IIId) comprising at least one first polypeptide
    selected from (I) or FP, and optionally comprising an additional
    polypeptide which enhances the immune response to the first polypeptide;
          (16) a pharmaceutical composition comprising (I), FP, (IIIc) or Ab;
```

selected from (I), (II), FP and Ab;

(18) identifying (I) or FP which induces an immune response effective to prevent or lessen the severity of Chlamydia infection in a mammal previously immunized with polypeptide, by immunizing a mouse with (I) or FP, and inoculating the immunized mouse with Chlamydia, where (I) or FP

(17) a diagnostic kit comprising instructions for use and a component

```
(19) an expression plasmid pCACPNM643 given in the specification;
           (20) a nucleic acid molecule comprising a sequence (S7); and
           (21) a peptide comprising a sequence (S8).
           gcgccggatcccagaqtcttqcagacgggg.
           (S8) is AlaLysTyrArgLysLysGlnGluAlaSerValLysLysTyrGln or
     TyrLeuPhePheProGlyTyrTyrThr.
          ACTIVITY - Antibacterial.
          MECHANISM OF ACTION - Vaccine (claimed); gene therapy.
          Groups of 7-9 week old male Balb/c mice (8-10 per group) were
     immunized intramuscularly (i.m.) and intranasally (i.n.) with plasmid DNA
     containing Chlamydia pneumoniae transmembrane protein gene. Saline or
     plasmid vector lacking an inserted Chlamydial gene was given to groups of
     control animals. At week 8, immunized mice were inoculated i.n. with 5 multiply 105 infection forming units (IFU) of C. pneumoniae strain AR39 to
     test their ability to limit the growth of a sublethal C. pneumoniae
     challenge. Lungs were taken from mice at day 9 post-challenge and
     immediately homogenized for analyzing the presence of Chlamydial
     inclusions using convalescent sera from rabbits infected with C.
     pneumoniae and metal-enhanced DAB as a peroxidase substrate. The results
     showed that mice immunized with pCACPNM643 had Chlamydial lung titers less
     than 60000 in 5/6 cases at day 9 (mean 37993), and values for control mice sham immunized with saline was 53100-315200 IFU/lung (mean 141593) at day
          USE - (I), (II), (III), PC and Ab are useful for preventing or
     treating Chlamydia infection. (I), (II) and Ab are useful for detecting
     Chlamydia infection, by assaying a body fluid of a mammal to be tested
     (claimed). (I) and (II) are useful as vaccines. The probes are used in
     diagnostic tests as capture or detection probes and in hybridization
     techniques, and primers are useful in amplification techniques for use in
     diagnostic methods. (I) is useful for detecting the presence of
     anti-Chlamydia antibodies in blood sample.
     Dwg.0/4
FS
     CPI EPI
     AB; DCN
FΑ
     CPI: B04-C01B; B04-C01C; B04-E01; B04-E03F; B04-E05; B04-E08; B04-F0100E;
MC
          B04-F10A; B04-G01; B04-N03A0E; B04-P01A; B11-C08;
          B11-C08E2; B11-C08E5; B12-K04A4; B12-K04E; B12-K04F; B14-A01A
          ; B14-S03; B14-S11B; D05-C12; D05-H07; D05-H09; D05-H12A;
          D05-H12D1; D05-H12E; D05-H14; D05-H17A6
     EPI: S03-E14H; S03-E14H4
L54 ANSWER 3 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
     2001-648558 [74] WPIX
AN
                         DNC C2001-191445
DNN N2001-484574
     Novel Chlamydia myosin heavy chain homolog polypeptide and polynucleotide
     for preventing, detecting and treating Chlamydia infections in mammals, in
     particular humans.
DC
     B04 D16 S03
     DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J (AVET) AVENTIS PASTEUR LTD; (DUNN-I) DUNN P; (MURD-I) MURDIN A
TN
     D; (OOME-I) OOMEN R P; (WANG-I) WANG J
CYC
    95
                     A2 20011011 (200174)* EN
PΤ
     WO 2001075113
                                                  83
                                                          C12N015-31
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TR TZ UG ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK
            DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ
            LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD
            SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
                     A 20011015 (200209)
A1 20020919 (200264)
     AU 2001048177
                                                          C12N015-31
     US 2002132994
                                                          C07H021-02
ADT WO 2001075113 A2 WO 2001-CA461 20010404; AU 2001048177 A AU 2001-48177
     20010404; US 2002132994 Al Provisional US 2000-194475P 20000404, US
     2001-824568 20010403
    AU 2001048177 A Based on WO 2001075113
FDT
PRAI US 2000-194475P
                           20000404; US 2001-824568
                                                             20010403
     ICM C07H021-02; C12N015-31
          A61K039-118; A61K039-40; A61K048-00; C07H021-04; C07K014-295; C07K016-12; C12N015-11; C12N015-62; C12Q001-68;
          G01N033-53; G01N033-68
AB
     WO 200175113 A UPAB: 20021031
     NOVELTY - An isolated myosin heavy chain homolog polypeptide (I) from
     Chlamydia, especially C. pneumoniae having a 254 residue amino acid sequence (S1), fully defined in the specification, its immunogenic
```

are identified;

fragment comprising at least 12 consecutive amino acids or a polypeptide which has been modified without loss of immunogenicity and which has 75 \$ sequence identity to (I), is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a nucleic acid molecule (II) comprising a nucleic acid sequence chosen from a sequence which encodes (I), a 965 base pair sequence (S2), fully defined in the specification, a sequence which encodes a polypeptide encoded by (S2), a sequence comprising 38 consecutive nucleotides from (II) and a sequence which encodes a polypeptide which is 75 % identical in amino acid sequence to the polypeptide encoded by (S2);
- (2) a nucleic acid molecule (III) comprising a nucleic acid sequence which is anti-sense to (II);
  - (3) a fusion protein (IV) comprising (I) and a second polypeptide;
- (4) a nucleic acid molecule  $(\bar{V})$  comprising a nucleic acid sequence which encodes (IV);
- (5) a nucleic acid molecule chosen from (II), (III) and (V) operatively linked to one or more expression control sequences;
  - (6) a vaccine (VI), comprising:
- (a) a vaccine vector and (II), (III) or (V), where each nucleic acid is capable of being expressed and the vaccine optionally comprises a second nucleic acid encoding and capable of expressing an additional polypeptide which enhances the immune response to the polypeptide expressed by the nucleic acids; or
- (b) (I) or (IV), optionally comprising an additional polypeptide which enhances the immune response to (I) or (IV);
  - (7) a unicellular host (VII) transformed with (II), (III) or (V);
- (8) an isolated nucleic acid probe of 5-100 nucleotides which hybridizes under stringent conditions to (II), its complement or anti-sense sequence;
- (9) an isolated primer of 10-40 nucleotides which hybridizes under stringent conditions to (II), its complement or anti-sense sequence;
  - (10) a polypeptide encoded by (II) or (V);
  - (11) producing (I) or (IV), comprising culturing (VII);
  - (12) an antibody (VIII) against (I) or (IV);
  - (13) a pharmaceutical composition (IX) comprising (II), (III), (V),
- (I), (VI) or (VIII);
- (14) a diagnostic kit comprising instructions for use and (II), (III), (V), (I), (IV) or (VIII);
- (15) identifying (I) or (IV) which induces an immune response effective to prevent or lessen the severity of Chlamydia infection in a mammal previously immunized with polypeptide, by immunizing a mouse with (I) or (IV) and inoculating the immunized mouse with Chlamydia, where (I) or (IV) which prevents or lessens the severity of Chlamydia infection in the immunized mouse compared to a non-immunized control mouse is identified;
- (16) expression plasmid pCACPNM559 containing the myosin heavy chain homolog gene, as shown in the specification;
  - (17) a nucleic acid molecule of sequence (S7); and
  - (18) a peptide having the sequence (S8).
- (S7) is ATAAGAATGCGGCCGCCACCATGCATGACGCACTTCTAAGCA or GCGCCGGATCCCTACAGCTGCGCGACGACGACG.
- (S8) is ArgValLysLysGluHisGlnLysGluLeu, LysMetAspGluPheAsnAlaLeuThr, TrpGlnGluSerGlnValAsnAlaGlnGluAsnSerThrAlaLysArgArgArgArgArgA, AlaLeuLeuGluGlnArgThrGluLeu or IleLeuTyrTrpGlnGluSerGlnVal.

ACTIVITY - Antibiotic.

MECHANISM OF ACTION - Vaccine.

The effect of C. pneumoniae myosin heavy chain homolog gene in protecting mice against an intranasal challenge of C. pneumoniae was studied. Strain AR-39 was used in Balb/c mice as a challenge infection model to examine the capacity of Chlamydia gene products delivered as naked DNA to elicit a protective response against a sublethal C. pneumoniae lung infection. Protective immunity was defined as an accelerated clearance of pulmonary infection. Groups of 7-9 week old male Balb/c mice were immunized intramuscularly (i.m.) plus intranasally (i.n.) with plasmid DNA containing the C. pneumoniae myosin heavy chain homolog gene (pCACPNM559). Saline or the plasmid vector lacking an inserted chlamydial gene was given to groups of control animals. For i.m. immunization, alternate left and right quadriceps were injected with 100 micro g of DNA in 50 micro l of phosphate buffered saline (PBS) on three occasions at 0, 3 and 6 weeks. For i.n. immunization, anesthetized mice were aspirated 50 micro 1 of PBS containing 50 micro g DNA on three occasions at 0, 3 and 6 weeks. At week 8, immunized mice were inoculated i.n. with 5 multiply 105 infection forming units (IFU) of C. pneumoniae, strain AR39 in 100 micro l of SPG buffer to test their ability to limit the growth of a sublethal C. pneumoniae challenge. Lungs were taken from

Graser 10/030313

Page 66

mice at day 9 post-challenge and immediately homogenized. Dilutions of the homogenate were assayed for the presence of infectious Chlamydia. The results showed that the mice immunized i.n. and i.m. with pCACPNM559 had chlamydial lung titers less than 49000 in 5 of 6 cases at day 9 and for control mice sham immunized with saline the value was 53100-315200 IFU/lung at day 9. USE - (I)-(V) and (VIII) are useful for detecting Chlamydia infection by assaying a body fluid of a mammal with the components. (VI) and (IX)

are useful for preventing and treating Chlamydia infection (claimed), in mammals, such as humans. The nucleic acid molecules are useful for producing (I), in the construction of vaccine vectors such as poxviruses, which are further useful for preventing and/or treating Chlamydia infection and in the construction of attenuated Chlamydia strains that can over-express the nucleic acid molecules or express it in a non-toxic, mutated form. (VI) is effective in preventing and/or treating Chlamydia infection for e.g. infection caused by C. trachomatis, C. psittaci, C. pneumoniae or C. pecorum. Probes which hybridize to (II) are useful in diagnostic tests, as capture of detection probes. (VIII) is useful in affinity chromatography for purifying (I) and in prophylactic or therapeutic passive immunization methods. Dwq.0/4

FS CPI EPI

AB; DCN FΑ

CPI: B04-C01B; B04-C01D; B04-C01G; B04-E01; B04-E03F; B04-E05; B04-E08; MC

B04-F10A; B04-G07; B04-N03A0E; B04-P01A; B11-C08; B11-C08E2; B11-C08E5; B12-K04A4; B12-K04E; B12-K04F; B14-A01A;

B14-S03; B14-S11B; D05-C12; D05-H07; D05-H09; D05-H12A;

D05-H12D1; D05-H12E; D05-H14; D05-H17A6

EPI: S03-E14H; S03-E14H4

L54 ANSWER 4 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN

2001-648557 [74] WPIX AN

DNN N2001-484573 DNC C2001-191444

Novel Chlamydia glutamate binding protein and polynucleotide for preventing, detecting and treating Chlamydia infections in mammals, in particular humans.

DC B04 D16 S03

DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J IN

(AVET) AVENTIS PASTEUR LTD; (DUNN-I) DUNN P; (MURD-I) MURDIN A PA D; (OOME-I) OOMEN R P; (WANG-I) WANG J

CYC 95

WO 2001075112 A2 20011011 (200174)\* EN 86 C12N015-31 PΙ RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD

SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW C12N015-31

AU 2001048176 A 20011015 (200209) US 2002094965 A1 20020718 (200254) A61K048-00

WO 2001075112 A2 WO 2001-CA460 20010404; AU 2001048176 A AU 2001-48176 20010404; US 2002094965 Al Provisional US 2000-194472P 20000404, US 2001-824206 20010403

FDT AU 2001048176 A Based on WO 2001075112

PRAI US 2000-194472P 20000404; US 2001-824206 20010403

ICM A61K048-00; C12N015-31

ICS A61K039-118; A61K039-40; C07H021-04; C07K014-295; C07K016-12; C07K019-00; C12N001-21; C12N015-62; C12N015-74;

C12Q001-68; G01N033-569 WO 200175112 A UPAB: 20011217

NOVELTY - An isolated glutamate binding protein (I) from Chlamydia, especially C. pneumoniae having a 250 residue amino acid sequence (S1), fully defined in the specification, its immunogenic fragment comprising at least 12 consecutive amino acids or a polypeptide which has been modified without loss of immunogenicity and which has 75 % sequence identity to (I), is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the

(1) a nucleic acid molecule (II) comprising a nucleic acid sequence chosen from a sequence which encodes (I), a 953 base pair sequence (S2) fully defined in the specification, a sequence which encodes a polypeptide encoded by (S2), a sequence comprising 38 consecutive nucleotides from (II) and a sequence which encodes a polypeptide which is 75 % identical in amino acid sequence to the polypeptide encoded by (S2);

(2) a nucleic acid molecule (III) comprising a nucleic acid sequence which is anti-sense to (II);

- (3) a fusion protein (IV) comprising (I) and a second polypeptide;(4) a nucleic acid molecule (V) comprising a nucleic acid sequence which encodes (IV);
- (5) a nucleic acid molecule chosen from (II), (III) and (V) operatively linked to one or more expression control sequences;
  - (6) a vaccine (VI), comprising:
- (a) a vaccine vector and any one of the above nucleic acids, where each nucleic acid is capable of being expressed and the vaccine optionally comprises a second nucleic acid encoding and capable of expressing an additional polypeptide which enhances the immune response to the polypeptide expressed by the nucleic acid; or
- (b) (I) or (IV), optionally comprising an additional polypeptide which enhances the immune response to (I) or (IV);
  - (7) a unicellular host (VII) transformed with (II), (III) or (V);(8) an isolated nucleic acid probe of 5-100 nucleotides which
- hybridizes under stringent conditions to (II), its complement or anti-sense sequence;
- (9) an isolated primer of 10-40 nucleotides which hybridizes under stringent conditions to (II), its complement or anti-sense sequence;
  - (10) a polypeptide encoded by (II) or (V);
  - (11) producing (I) or (IV), comprising culturing (VII);
  - (12) an antibody (VIII) against (I) or (IV);
- (13) a pharmaceutical composition (IX) comprising (II), (III), (V),
  (I), (VI) or (VIII);
- (14) a diagnostic kit comprising instructions for use and (II), (III), (V), (I), (IV) or (VIII);
- (15) identifying (I) or (IV) which induces an immune response effective to prevent or lessen the severity of Chlamydia infection in a mammal previously immunized with polypeptide, by immunizing a mouse with (I) or (IV) and inoculating the immunized mouse with Chlamydia, where (I) or (IV) which prevents or lessens the severity of Chlamydia infection in the immunized mouse compared to a non-immunized control mouse is identified;
- (16) expression plasmid pCACPNM653 containing the glutamate binding protein gene;
  - (17) a nucleic acid molecule of sequence (S7); and
  - (18) a peptide having the sequence (S8).
- (S7) is ATAAGAATGCGGCCGCCACCATGAAGATAAAATTTTCTTGGAAGG or GCGCCGGATCCCGGGAAGACGATACCGCTGTTTT. (S8) is GluAsnLeuAspAspLysLysThrGlnGly, LysThrArgArgSerGlyLysTyrAspAlaIleLysGlnArgTyrArgLeuPro, AlaLeuLeuAlaProValIleGluVal or PheLeuAsnAspLeuValSerGluIle.

ACTIVITY - Antibiotic.

MECHANISM OF ACTION - Vaccine.

The effect of C. pneumoniae glutamate binding protein gene in protecting mice against an intranasal challenge of C. pneumoniae was studied. Strain AR-39 Grayston et al (1990) Journal of Infectious Diseases 161:618-625 was used in Balb/c mice as a challenge infection model to examine the capacity of Chlamydia gene products delivered as naked DNA to elicit a protective response against a sublethal C. pneumoniae lung infection. Protective immunity was defined as an accelerated clearance of pulmonary infection. Groups of 7-9 week old male Balb/c mice were immunized intramuscularly (i.m.) plus intranasally (i.n.) with plasmid DNA containing the C. pneumoniae glutamate binding protein gene (pCACPNM653). Saline or the plasmid vector lacking an inserted chlamydial gene was given to groups of control animals. For i.m. immunization, alternate left and right quadriceps were injected with 100 micro g of DNA in 50 micro l of phosphate buffered saline (PBS) on three occasions at 0, 3 and 6 weeks. For i.n. immunization, anesthetized mice were aspirated 50 micro l of PBS containing 50 micro g DNA on three occasions at  $\bar{\text{0}}$ , 3 and 6 weeks. At week 8, immunized mice were inoculated i.n. with 5 multiply 105 infection forming units (IFU) of C. pneumoniae, strain AR39 in 100 micro 1 of SPG buffer to test their ability to limit the growth of a sublethal C. pneumoniae challenge. Lungs were taken from mice at day 9 post-challenge and immediately homogenized. Dilutions of the homogenate were assayed for the presence of infectious Chlamydia. The results showed that the mice immunized i.n. and i.m. with pCACPNM653 had chlamydial lung titers less than 60000 in 4 of 6 cases at day 9 and for control mice sham immunized with saline the value was 53100-315200 IFU/lung at day 9.

USE - (I)-(V) and (VIII) are useful for detecting Chlamydia infection by assaying a body fluid of a mammal with the components (claimed). (VI) and (IX) are useful for preventing or treating Chlamydia infection (claimed), in mammals, such as humans. The nucleic acid molecules are useful for producing (I), in the construction of vaccine vectors such as poxviruses, which are further useful for preventing and/or treating Chlamydia infection and in the construction of attenuated Chlamydia strains that can over-express the nucleic acid molecules or express it in

a non-toxic, mutated form. (VI) is effective in preventing and/or treating Chlamydia infection for e.g. infection caused by C. trachomatis, C. psittaci, C. pneumoniae or C. pecorum. Probes which hybridize to (II) are useful in diagnostic tests, as capture of detection probes. (VIII) is useful in affinity chromatography for purifying (I) and in prophylactic or therapeutic passive immunization methods. Dwq.0/4 FS CPI EPI AB; DCN CPI: B04-C01B; B04-C01D; B04-C01G; B04-E01; B04-E03F; B04-E05; B04-E08; B04-F0100E; B04-F10A; B04-N03A0E; B11-C07A; B11-C08E2; B11-C08E5; B12-K04A4; B12-K04E; B12-K04F; B14-A01A; B14-S03; B14-S11B; D05-C12; D05-H07; D05-H09; D05-H11; D05-H12A; D05-H12D1; D05-H12E; D05-H14; D05-H17A6 EPI: S03-E14H4 L54 ANSWER 5 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN 2001-648556 [74] WPIX DNC C2001-191443 DNN N2001-484572 Novel isolated myosin heavy chain polypeptide from Chlamydia pneumoniae and polynucleotides encoding them, useful for treating or preventing Chlamydia infection in mammals. DC B04 D16 S03 DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J TN (AVET) AVENTIS PASTEUR LTD; (DUNN-I) DUNN P; (MURD-I) MURDIN A PA D; (OOME-I) OOMEN R P; (WANG-I) WANG J CYC WO 2001075111 A2 20011011 (200174)\* EN 83 C12N015-31 PΤ RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2001048172 A 20011015 (200209) C12N015-31 US 2003100706 Al 20030529 (200337) A61K039-02 WO 2001075111 A2 WO 2001-CA456 20010404; AU 2001048172 A AU 2001-48172 ADT 20010404; US 2003100706 A1 Provisional US 2000-194471P 20000404, US 2001-824584 20010403 FDT AU 2001048172 A Based on WO 2001075111 20000404; US 2001-824584 PRAT US 2000-194471P 20010403 ICM A61K039-02; C12N015-31 A61K039-118; A61K039-40; A61K048-00; C07H021-04; C07K014-195; C07K014-295; C07K016-12; C07K019-00; C12N001-21; C12N015-62; C12N015-74; C12Q001-68; G01N033-569 WO 200175111 A UPAB: 20011217 NOVELTY - An isolated myosin heavy chain polypeptide (I) from Chlamydia pneumoniae, comprising 168 residue amino acid sequence (S2), fully defined in the specification, an immunogenic fragment having 12 consecutive amino acids of (S2), or a sequence of (S2) or its fragment which has been modified without loss of immunogenicity and having 75 % identity to above mentioned polypeptide sequences, is new. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following: (1) a nucleic acid molecule (II) comprising a nucleic acid sequence which encodes (I), comprising: (a) a 707 nucleotide sequence (S1), fully defined in specification; (b) a sequence which encodes a polypeptide encoded by (S1); (c) a sequence comprising at least 38 consecutive nucleotides of (a) or (b), or a sequence which encodes a polypeptide that is 75 % identical in amino acid sequence to polypeptide encoded by (S1); (2) a nucleic acid molecule (III) comprising a nucleic acid sequence which is antisense to (II); (3) a nucleic acid molecule (IV) comprising a nucleic acid sequence which encodes fusion protein that comprises a polypeptide encoded by (II) and a second polypeptide; (4) a nucleic acid molecule ((I)-(IV)) operatively linked to one or more expression control sequences; (5) a vaccine (V) comprising a vaccine vector and (II); (6) a vaccine (VI) comprising a vaccine vector and at least one first nucleic acid encoding a fusion protein that comprises a polypeptide encoded by (S1), a polypeptide encoded by a nucleic acid comprising at least 38 consecutive nucleotides from (S1), a polypeptide which is 75 % identical to the amino acid sequence to the polypeptide encoded by (S1), or is (I); and

(7) a vaccine (VII) comprising (II), (III), or (V) operatively linked

to expression control sequences, as first nucleic acid and a vaccine vector, the vaccine optionally comprises a second nucleic acid encoding an additional polypeptide which enhances the immune response to the polypeptide expressed by the first;

- (8) a unicellular host (VIII) transformed with a nucleic acid molecule ((II)-(IV)) operatively linked to expression control sequences;
- (9) an isolated nucleic acid probe (IX) of 5-100 nucleotides which hybridizes under stringent conditions to (S1);
- (10) an isolated primer (X) of 10-40 nucleotides which hybridizes under stringent conditions to (S1);
- (11) a polypeptide encoded by (II), (III), (IV) or a nucleic acid molecule ((II)-(IV)) operatively linked to expression control sequences;
  - (12) a fusion protein (XI) comprising (I) and a second polypeptide;
  - (13) preparation of (I) or (XI);
  - (14) an antibody (XII) against (I) or (XI);
- (15) a vaccine (XIII) comprising at least one first polypeptide (FP1) encoded by (S1), a polypeptide encoded by a nucleic acid comprising at least 38 consecutive nucleotides of (S1), a polypeptide which is 75 % identical to the amino acid sequence to the polypeptide encoded by (S1), or is (I), where the vaccine optionally comprises an additional polypeptide which enhances the immune response to FP1;
- (16) a vaccine (XIV) comprising a fusion protein which comprises FP1 and a second polypeptide, where the vaccine optionally comprises an additional polypeptide which enhances the immune response to FP1;
- (17) a vaccine (XV) comprising (I) or (XI) as the first polypeptide, and an additional polypeptide which enhances the immune response to the first polypeptide;
- (18) a diagnostic kit comprising instructions for use and a component
  (II), (III), (V) operatively linked to expression control sequences, (I),
  (XI) or (XII);
- (19) identifying (I) or (XI) which prevents or lessens the severity of Chlamydia infection in a mammal previously immunized with polypeptide involves immunizing a mouse with the polypeptide or fusion protein and inoculating the immunized mouse with Chlamydia;
  - (20) expression plasmid pCACPNM760;
  - (21) a nucleic acid molecule having a sequence (S7); and
  - (22) a peptide having a sequence (S8).
- (S7) is ataagaatgcggccgccaccatggcaaaatatccactagagcc or gcgccggatcccgcttccccctgattcacg.
- (S8) is LysArgArgLysGluGluIuLysThrArgLeuHisLysGluGluTrpMet, LeuArgGlnLysLysArgGluSerGlyGlySer or GlnLeuSerGluGluGluGluIuLysVal. ACTIVITY - Antibiotic.

MECHANISM OF ACTION - Vaccine.

Strain AR-39 was used in Balb/c mice as a challenge infection model to examine the capacity of Chlamydia gene products delivered as naked DNA to elicit a protective response against sublethal C. pneumoniae lung infections. Groups of 7-9 week old male Balb/c mice were immunized intramuscularly (i.m.) plus intranasally (i.n.) with plasmid DNA containing the C. pneumoniae myosin heavy chain gene. For i.m. immunization, alternate left and right quadriceps were injected with 100 micro g of DNA. For i.n. immunization, anesthetized mice were aspirated 50 micro I of phosphate buffered saline (PBS) containing 50 micro g DNA on three occasions at 0, 3 and 6 weeks. At week 8, immunized mice were inoculated i.n. with 5 multiply 105 infection forming units (IFU) of C. pneumoniae, strain AR39 in 100 micro l of SPG buffer to test their ability to limit the growth of a sublethal C. pneumoniae challenge. Lungs were taken from mice at day 9 post-challenge and homogenized in SPG buffer. Dilutions of the homogenate were assayed for Chlamydia by inoculation onto monolayers of susceptible cells. The inoculum was centrifuged onto the cells, then the cells were incubated for three days at 35 deg. C in the presence of 1 micro g/ml cycloheximide. After incubation the monolayers were fixed and stained using convalescent sera from rabbits infected with C. pneumoniae. Results showed that mice immunized with i.n. and i.m. with pCACPNM760 had chlamydial lung titers less than 40000 in 3 of 6 cases at day 9, whereas the range of values for control mice sham immunized with saline was 20800-323300 IFU/lung at day 9.

USE - (II), (III), (IV) or a nucleic acid molecule ((II), (III), (V)) operatively linked to expression control sequences, the vaccines and pharmaceutical compositions are useful for treating or preventing Chlamydia infection. (II), (III), (IV) or a nucleic acid molecule ((II)-(IV)) operatively linked to expression control sequences, (I), (XI) or (XII) is also useful for detecting Chlamydia infection. (All claimed). (I) is useful for detecting the presence of anti-Chlamydia antibodies in a biological sample. (II) is useful for producing (I), for constructing vaccine vectors, and as a vaccine agent, or in the construction of attenuated Chlamydia strains that can overexpress (II). (IX) is useful as

capture or detection probe. (IX) and (X) are useful for detecting and/or identifying the presence of Chlamydia in a biological material. (XII) is useful for purifying (I) by antibody-based affinity chromatography. (XII) can also be used in therapeutic and prophylactic passive immunization methods. (XII) used for detecting Chlamydia in biological sample. Dwg.0/4 CPI EPI FS AB; DCN FA MC CPI: B04-C01B; B04-C01C; B04-C01D; B04-C01G; B04-E01; B04-E03F; B04-E05; B04-E08; B04-F0100E; B04-F10A; B04-G07; B04-N03A0E; B11-C08; B11-C08E2; B11-C08E5; B12-K04A4; B12-K04E; B12-K04F; B14-A01A ; B14-S03; B14-S11B; D05-C12; D05-H07; D05-H09; D05-H12A; D05-H12D1; D05-H12E; D05-H14; D05-H17A6 EPI: S03-E14H4 L54 ANSWER 6 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN 2001-343797 [36] WPIX DNC C2001-106482 A Chlamydia polypeptide, an amino acid transporter gene, for the treatment TΙ and prevention of Chlamydia infection. DC B04 C06 D16 DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J IN (AVET) AVENTIS PASTEUR LTD PA CYC 94 WO 2001036457 A2 20010525 (200136)\* EN PΙ 81 C07K014-00 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2001013757 A 20010530 (200152) C07K014-00 WO 2001036457 A2 WO 2000-CA1346 20001110; AU 2001013757 A AU 2001-13757 ADT 20001110 AU 2001013757 A Based on WO 2001036457 PRAI US 1999-165615P 19991115 ICM C07K014-00 TC WO 200136457 A UPAB: 20010628 NOVELTY - A Chlamydia polypeptide which is encoded by (I), a 468 amino acid (aa) sequence, given in the specification, is new DETAILED DESCRIPTION - The polypeptide may also be a fusion protein comprising (I) and an additional polypeptide. INDEPENDENT CLAIMS are included for the following: (1) a nucleic acid molecule which encodes a polypeptide, a C. pneumoniae, an amino acid transporter gene comprising: (i) a 1607 base pair (bp) nucleic acid sequence defined in the specification; (ii) an immunogenic fragment comprising at least 12 aa from a polypeptide encoded by (a); and (iii) a polypeptide of (a) or (b) which has been modified to improve its immunogenicity, where the peptide is at least 75% identical in aa sequence to (a) or (b). (2) a nucleic acid sequence selected from: (i) 1564 bp sequence; a sequence including (a); (ii) a sequence which encodes a polypeptide encoded by (i); (iii) a sequence comprising at least 38 consecutive nucleotides from (i) or (ii); (iv) a sequence which encodes a polypeptide which is at least 75% identical in aa sequence to the polypeptide encoded by (i); (3) a nucleic acid molecule comprising a nucleic acid sequence which is antisense to (1); (4) a nucleic acid molecule comprising a sequence encoding a fused protein which is encoding a nucleic acid (1) and an additional polypeptide; (5) a vaccine comprising a nucleic acid of (1) and a vaccine vector where each nucleic acid is expressed as a polypeptide. The vaccine optionally comprises a second nucleic acid encoding an additional polypeptide which enhances the immune response to the polypeptide is expressed by the first;

Search done by Noble Jarrell

(6) a unicellular host transformed with the nucleic acid molecule
 (4);
 (7) a nucleic acid probe of 5 to 100 nucleotides which hybridizes under stringent conditions to the nucleic acid molecule (1) or to its

(8) a primer of 10 to 40 nucleotides which hybridizes under stringent conditions to the nucleic acid molecule of (1) or to its homolog or

homolog or complementary anti-sense sequence;

```
complementary anti-sense sequence;
          (9) the production of the polypeptide (I) comprising the culturing of
          (10) an antibody against polypeptide(s) of the invention;
          (11) a vaccine comprising at least one first polypeptide of the
     invention, and optionally a second polypeptide which enhances the immune
     response to the first;
          (12) the treatment or prevention of Chlamydia infection using:
          (i) a nucleic acid of (1-4);
          (ii) a vaccine of (5) or (11);
          (iii) a polypeptide of the invention; and/or
     (iv) (11);
          (13) the detection of Chlamydia comprising the step of assaying a
     body fluid of a mammal with a component selected from 12 (i), (iii) and/or
     (11);
          (14) a diagnostic kit comprising instructions for use and 12 (i),
     (iii) or (11);
          (15) the identification of a polypeptide of the invention which
     induces a response effective to prevent or lessen the extent of Chlamydia
     infection in a mammal previously immunized with a polypeptide comprising:
          (i) immunizing a mouse with the polypeptide; and
          (ii) innoculuating the immunized mouse with Chlamydia; where the
     polypeptide which prevents or lessens the severity of Chlamydia infection
     in the immunized mouse compared to a non-immunized control mouse
          (16) expression plasmid pCABk297.
          ACTIVITY - Antibacterial.
          MECHANISM OF ACTION - Vaccine. Gene therapy.
          USE - The polypeptide, an amino acid transporter is useful for the
     treatment, prevention and diagnosis of Chlamydia infection, preferably
     Chlamydia pneumoniae infection (claimed), in human and veterinary
     applications.
          ADVANTAGE - A protective vaccine against Chlamydia pneumoniae is
     useful to prevent infection which leads to chronic bronchitis and
     sinusitis. There is also a correlation between infection and
     atherosclerosis, with epidemiological studies showing connections between
     the incidence of heart attack, coronary artery and carotid artery disease with organisms being detected in the fatty streaks of the coronary,
     carotid, peripheral arteries and aorta. The infection may also be linked
     with the high incidence of lower respiratory tract infections and
     mortality in infants and children in tropical regions of the world. The
     preventative vaccine reduces the need for antibiotic treatment.
     Dwg.0/3
    CPI
    AB; DCN
     CPI: B04-C01G; B04-E02F; B04-E03F; B04-E05; B04-E06; B04-E08;
          B04-G01; B04-N03A; B11-C08E; B12-K04A4; B12-K04F;
          B14-A01A; C04-C01G; C04-E02F; C04-E03F; C04-E05; C04-E06;
          C04-E08; C04-G01; C04-N03A; C11-C08E; C12-K04A4; C12-K04F;
          C14-A01A; D05-C11; D05-H07; D05-H11; D05-H12D; D05-H12D1; D05-H12E; D05-H14; D05-H17A; D05-H17B
L54 ANSWER 7 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
     2001-343796 [36]
                        WPTX
DNC C2001-106481
     A Chlamydia polypeptide, OppB, for the treatment and prevention of
     Chlamydia infection.
    DUNN, P; MURDIN, A D; OOMEN, R P.; WANG, J
     (AVET) AVENTIS PASTEUR LTD
   94
     WO 2001036456
                     A2 20010525 (200136) * EN
                                                75
                                                       C07K014-00
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TR TZ UG ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM
            DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC
            LK LR'LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE
            SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
     AU 2001013756 A 20010530 (200152)
                                                        C07K014-00
    WO 2001036456 A2 WO 2000-CA1345 20001110; AU 2001013756 A AU 2001-13756
     20001110
   AU 2001013756 A Based on WO 2001036456
PRAI US 1999-164918P
                          19991115
    ICM C07K014-00
    WO 200136456 A UPAB: 20010628
     NOVELTY - A Chlamydia polypeptide which is encoded by (I) a 314 amino acid
     (aa) sequence, given in the specification, is new.
```

FS

FΑ

MC

ΔN

TI

IN

PA CYC

ADT

IC

DETAILED DESCRIPTION - The polypeptide may also be a fusion protein comprising (I) and an additional polypeptide.

- INDEPENDENT CLAIMS are included for the following:
- (1) a nucleic acid molecule which encodes a polypeptide, a C. pneumoniae, OppB gene comprising:
- (i) a 1145 base pair (bp) nucleic acid sequence defined in the specification;
- (ii) an immunogenic fragment comprising at least 12 aa from a polypeptide encoded by (a); and
- (iii) a polypeptide of (a) or (b) which has been modified to improve its immunogenicity, where the peptide is at least 75% identical in aa sequence to (a) or (b).
  - (2) a nucleic acid sequence selected from
  - (i) a sequence including (a);
  - (ii) a sequence which encodes a polypeptide encoded by (i);
- (iii) a sequence comprising at least 38 consecutive nucleotides from (i) or (ii);
- (iv) a sequence which encodes a polypeptide which is at least 75% identical in aa sequence to the polypeptide encoded by (i);
- (3) a nucleic acid molecule comprising a nucleic acid sequence which is antisense to (1);
- (4) a nucleic acid molecule comprising a sequence encoding a fused protein which is encoding a nucleic acid (1) and an additional polypeptide;
- (5) a vaccine comprising a nucleic acid of (1) and a vaccine vector where each nucleic acid is expressed as a polypeptide. The vaccine optionally comprises a second nucleic acid encoding an additional polypeptide which enhances the immune response to the polypeptide is expressed by the first;
- (6) a unicellular host transformed with the nucleic acid molecule
- (7) a nucleic acid probe of 5 to 100 nucleotides which hybridizes under stringent conditions to the nucleic acid molecule (a) or to its homolog or complementary anti-sense sequence;
- (8) a primer of 10 to 40 nucleotides which hybridizes under stringent conditions to the nucleic acid molecule of (a) or to its homolog or complementary anti-sense sequence;
- (9) the production of the polypeptide (I) comprising the culturing of (6):
  - (10) antibody against polypeptide(s) of the invention;
- (11) a vaccine comprising at least one first polypeptide of the invention, and optionally a second polypeptide which enhances the immune response to the first;
  - (12) the treatment of Chlamydia infection using:
  - (i) a nucleic acid of (1-4);
  - (ii) a vaccine of (5) or (12);
  - (iii) a polypeptide of the invention; and/or
- (iv) (10);
- (13) the detection of Chlamydia comprising the step of assaying a body fluid of a mammal with a component selected from 12 (i), (iii) and/or (iv);
- (14) a diagnostic kit comprising instructions for use and 12 (i),
  (iii) or (iv);
- (15) the identification of a polypeptide of the invention which induces a response effective to prevent or lessen the extent of Chlamydia infection in a mammal previously immunized with a polypeptide comprising:
  - (i) immunizing a mouse with the polypeptide; and
  - (ii) innoculuating the immunized mouse with Chlamydia;
  - (16) the expression plasmid pCAI434.

ACTIVITY - Antibacterial.

MECHANISM OF ACTION - Vaccine. Groups of 7-9 week old male Balb/c mice (n = 8-10 / group) were immunized intramuscularly (i.m) and intranasally (i.n.) with plasmid DNA containing the Chlamydia pneumonia OppB gene. Saline or the plasmid vector without the insert was given to the control animals. For i.m immunization, alternate left and right quadriceps were injected with 100 micro g of DNA in phosphate buffered saline (PBS) at three timepoints, 0, 3 and 6 weeks. For i.n immunization, anaesthetized mice were aspirated with 50 micro g of DNA in PBS at the three timepoints. A 8 weeks immunized mice were innoculated i.n with 5 X 105 IFU of Chlamydia pneumonia, strain AR39 in 100 micro l of SPG buffer. Lungs were taken from mice at day 9 post challenge, homogenised and the homegenate examined for the presence of Chlamydial inclusions. The mean bacterial load (inclusion forming units per lung) was 83378.6 for the saline control; 77000 for pCAI1021 (p = 0.7671); and 27450 for pCAI434 (p = 0.0028), where pCAI1021 and pCAI434 are active constructs.

USE - The polypeptide of the invention is useful for the treatment,

prevention and diagnosis of Chlamydia infection (claimed), preferably Chlamydia pneumonia infection, In human and veterinary applications. ADVANTAGE - A protective vaccine against Chlamydia pneumonia is useful to prevent infection which leads to chronic bronchitis and sinusitis. There is also a correlation between infection and atherosclerosis, with epidemiological studies showing connections between the incidence of heart attack, coronary artery and carotid artery disease with organisms being detected in the fatty streaks of the coronary, carotid, peripheral arteries and aorta. Dwg.0/4 CPI FS FA AB; DCN MC CPI: B04-C01G; B04-E02F; B04-E03F; B04-E05; B04-E06; B04-E08; B04-F0100E; B04-G01; B04-N03A; B11-C08E; B12-K04A4; B14-A01A; C04-C01G; C04-E02F; C04-E03F; C04-E05; C04-E06; C04-E08; C04-F01; C04-G01; C04-N03A; C11-C08E; C12-K04A4; C12-K04F; D05-C11; D05-H07; D05-H09; D05-H11; D05-H12A; D05-H12B; D05-H12D1; D05-H12D2; D05-H12E; D05-H14; D05-H17A; D05-H17B L54 ANSWER 8 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN 2001-328102 [34] WPIX AN DNN N2001-236077 DNC C2001-100610 New 1pxB polypeptides useful for treating, preventing or diagnosing Chlamydia infections, particularly infections caused by Chlamydia pneumonia, e.g. bronchitis, cough, asthma. B04 D16 S03 DC IN DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J (AVET) AVENTIS PASTEUR LTD PΑ CYC 94 WO 2001021810 A1 20010329 (200134)\* EN 80 C12N015-54 PΙ RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2000073982 A 20010424 (200141) C12N015-54 ADT WO 2001021810 A1 WO 2000-CA1085 20000915; AU 2000073982 A AU 2000-73982 20000915 AU 2000073982 A Based on WO 2001021810 19990917 PRAI US 1999-154461P ICM C12N015-54 ICS A61K031-711; A61K038-45; A61K039-40; C07K016-40; C12N009-10; C12N015-62; C12N015-85; G01N033-53 WO 200121810 A UPAB: 20011217 NOVELTY - A novel polypeptide (I) comprises: (A) a fully defined sequence (IIa) of 604 amino acids (aa) given in the specification; (B) an immunogenic fragment (IIb) comprising at least 12 consecutive aa from (IIa); (C) (IIa) or (IIb) which has been modified to improve its immunogenicity and is at least 75% identical to (IIa) or (IIb); (D) a sequence encoded by a sequence antisense to those in (A) - (C); (E) a polypeptide of (A) - (C) and an additional polypeptide. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the (1) a nucleic acid (II) encoding a polypeptide comprising: (a) a fully defined sequence (IIa) of 604 amino acids (aa) given in the specification; (b) an immunogenic fragment (IIb) comprising at least 12 consecutive aa from (IIa); or (c) (IIa) or (IIb) which has been modified to improve its immunogenicity and is at least 75% identical to (IIa) or (IIb), is new. (II) has a sequence of 2023 base pairs (bp) fully defined in the specification, or at least 38 consecutive nucleotides (nt) of this sequence. (2) a nucleic acid (III) comprising a sequence antisense to (II); (3) a nucleic acid (IV) encoding a fusion protein comprising a polypeptide encoded by (II) and an additional polypeptide; (4) vaccines (V) comprising: (a) at least one (II) a vaccine vector, and optionally a second nucleic acid encoding an additional polypeptide that enhances the immune response to the polypeptide expressed by the first nucleic acid; or (b) at least one (I) and optionally a second polypeptide that enhances the immune response to the first polypeptide;

Graser 10/030313

Page 74

(5) a unicellular host (VI) transformed with (II);

- (6) a nucleic acid probe (VIIa) of 5-100 nt or a primer (VIIb) of 10-40 nt, which hybridizes under stringent conditions to a 2023-bp sequence, or its homologue, complement, or antisense sequence;
  - (7) producing (I) by culturing (VI);
  - (8) an antibody (VIII) immunospecific for (I);
- (9) preventing or treating (M1) Chlamydia infection using the nucleic acids, vaccines, pharmaceutical composition, polypeptides or antibodies of the invention:
- (10) detecting (M2) Chlamydia infection by assaying a body fluid of a mammal with the nucleic acids, polypeptides or antibody of the invention;
- (11) a diagnostic kit (IX) comprising instructions for use and the nucleic acids, polypeptides or antibodies of the invention;
- (12) identifying (M3) a (I) that induces immune response effective to prevent or lessen the severity of Chlamydia infection in a mammal previously immunized with the polypeptide by:
  - (a) immunizing a mouse with the polypeptide; and
- (b) inoculating the immunized mouse with Chlamydia, where the polypeptide prevents or lessens the severity of Chlamydia infection in the immunized mouse compared to a non-immunized;
  - (13) expression plasmid (X), pCABk1043;
- (14) a nucleic acid (XI) with a 45 or 34 bp sequence given in the specification; and
  - (15) polypeptide 1pxB (XII) from Chlamydia.

ACTIVITY - Antimicrobial.

MECHANISM OF ACTION - Vaccine. 7-9 week old male Balb/c mice were immunized intramuscularly plus intranasally with plasmid DNA containing the coding sequence of C. pneumonia 1pxB or plasmid vector lacking an inserted chlamydial gene. Immunization was given at 0.3 and 6 weeks, and at week 8, mice were inoculated with 5 multiply 105 IFU of C. pneumoniae strain AR39.9 days post-challenge, lungs were taken and homogenized in SPG buffer. Dilutions of homogenate were assayed for the presence of infectious chlamydia by inoculation onto monolayers of susceptible cells. Cells were incubated for 3 days, and monolayers were fixed with formalin and methanol, and immunoperoxidase stained for the presence of chlamydial inclusions using convalescent sera from rabbits infected with C. pneumoniae. Mice immunized with pCABk1043 had chlamydial lung titers less than 37,000 in 5 of 6 cases at day 9, while the range of values for the controls was 13,600-458,100 IFU/lung.

USE - (I), (II), (V) and (VIII) are useful as pharmaceutical compositions (claimed). The nucleic acids encoding the Chlamydia 1pxB polypeptides are useful as a vaccine in preventing, treating or diagnosing Chlamydia infections, particularly those caused by C. pneumoniae, including respiratory diseases, e.g. cough, sore throat, bronchitis, asthma. The polynucleotides, including DNA or RNA may be used in producing the encoded polypeptide in a recombinant host system, in the construction of vaccine vectors such as pox viruses, as vaccine agent, and in constructing attenuated Chlamydia strains that can over-express a polynucleotide or express it in a non-toxic mutated form. The polypeptides may also be used as diagnostic reagent for detecting the presence of anti-Chlamydia antibodies, and in the preparation of a medicament for treating or preventing Chlamydia infection.

Dwg.0/4 FS CPI EPI

FA AB; DCN

MC CPI: B04-C01G; B04-E03F; B04-E05; B04-E06; B04-E08; B04-F01; B04-G07;

B04-N03A; B11-C07A; B11-C08E2; B11-C08E5; B12-K04A4; B12-K04F;

B14-A01A; B14-S11B; D05-C11; D05-H04; D05-H07;

D05-H08; D05-H11; D05-H12A; D05-H12C; D05-H12D1; D05-H12D2; D05-H12E;

D05-H14; D05-H17A5; D05-H17C

EPI: S03-E14H4

L54 ANSWER 9 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN

AN 2001-328101 [34] WPIX

DNN N2001-236076 DNC C2001-100609

TI New general secretion pathway protein E polypeptides and nucleic acids encoding the polypeptides useful for treating, preventing or diagnosing Chlamydia infections, particularly infections caused by Chlamydia pneumoniae.

DC B04 D16 S03

IN DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J

PA (AVET) AVENTIS PASTEUR LTD

CYC 94

WO 2001021805 A1 20010329 (200134)\* EN 79 C12N015-31
RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE

SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2000073986 A 20010424 (200141) C12N015-31

ADT WO 2001021805 A1 WO 2000-CA1089 20000915; AU 2000073986 A AU 2000-73986 20000915

FDT AU 2000073986 A Based on WO 2001021805

PRAI US 1999-154595P 19990917

IC ICM C12N015-31

ICS A61K031-711; A61K039-118; A61K039-40;

C07K014-295; C07K016-12; C12N015-62; C12N015-85; G01N033-53

AB WO 200121805 A UPAB: 20010620

NOVELTY - A nucleic acid (I) encoding a polypeptide comprising:

- (a) a fully defined sequence of 496 amino acids given in the specification;
- (b) an immunogenic fragment comprising at least 12 consecutive amino acids from (a); or
- (c) (a) or (b) which has been modified to improve its immunogenicity and which is at least 75% identical to (a) or (b), is new.

DETAILED DESCRIPTION - The nucleic acid (I) has a sequence of 1691 bp fully defined in the specification, or has at least 38 consecutive nucleotides of this sequence.

INDEPENDENT CLAIMS are also included for the following:

- a nucleic acid comprising a sequence antisense to (I);
- (2) a nucleic acid encoding a fusion protein comprising a polypeptide encoded by (I) and an additional polypeptide;
- (3) vaccines comprising at least one first nucleic acid expressed as a polypeptide, a vaccine vector, and optionally a second nucleic acid encoding an additional polypeptide that enhances the immune response to the polypeptide expressed by the first nucleic acid;
  - (4) a unicellular host transformed with the nucleic acid;
- (5) a nucleic acid probe of 5-100 nucleotides or a primer of 10-40 nucleotides, which hybridizes under stringent conditions to a 1691-bp sequence, or its homologue, complement, or antisense sequence;
  - (6) a polypeptide encoded by the nucleic acids;
- (7) vaccines comprising at least one first polypeptide and optionally a second polypeptide that enhances the immune response to the first polypeptide;
- (8) a fusion polypeptide comprising a polypeptide of (6) and an additional polypeptide;
- (9) a method of producing a polypeptide of (6) by culturing a unicellular host of (4);
  - (10) an antibody against the polypeptide of (6);
- (11) pharmaceutical compositions comprising a polypeptide or an antibody;
- (12) a method of preventing or treating Chlamydia infection using the above nucleic acids, vaccines, pharmaceutical composition, polypeptides or antibodies:
- (13) a method of detecting Chlamydia infection by assaying a body fluid of a mammal with the above nucleic acids, polypeptides or antibody;
- (14) a diagnostic kit comprising instructions for use and the above nucleic acids, polypeptides or antibodies;
- (15) a method for identifying a polypeptide that induces immune response effective to prevent or lessen the severity of Chlamydia infection in a mammal previously immunized with the polypeptide by:
  - (a) immunizing a mouse with the polypeptide; and
- (b) inoculating the immunized mouse with Chlamydia, where the polypeptide prevents or lessens the severity of Chlamydia infection in the immunized mouse compared to a non-immunized;
  - (16) expression plasmid pCAI284;
  - (17) the nucleic acid
  - (I) ATAAGAATGC GGCCGCCACC ATGGCTGCTA GTATTTTAT;
  - (II) CCCCAAGCTT CATCACAGCG CTTGGTAAC.
  - (18) having a 39 or 29 bp sequence given in the specification; and
- (19) general secretion pathway protein E from Chlamydia, preferably C. pneumoniae.

ACTIVITY - Antimicrobial.

MECHANISM OF ACTION - Vaccine. 7-9 week old male Balb/c mice were immunized intramuscularly plus intranasally with plasmid DNA containing the coding sequence of C. pneumoniae general secretion pathway protein E or plasmid vector lacking an inserted chlamydial gene. Immunization was given at 0.3 and 6 weeks, and at week 8, mice were inoculated with 5 multiply 105 IFU of C. pneumoniae strain AR39. 9 days post-challenge, lungs were taken and homogenized in SPG buffer. Dilutions of homogenate were assayed for the presence of infectious chlamydia by inoculation onto

Graser 10/030313 monolayers of susceptible cells. Cells were incubated for 3 days, and monolayers were fixed with formalin and methanol, and imunoperoxidase stained for the presence of chlamydial inclusions using convalescent sera from rabbits infected with C. pneumoniae. Mice immunized with pCAI284 had chlamydial lung titers less than 50,000 in 5 of 6 cases at day 9, while the range of values for the controls was 18,200-247,100 IFU/lung. USE - The nucleic acids encoding the Chlamydia general secretion pathway protein E polypeptides are useful as a vaccine in preventing, treating or diagnosing Chlamydia infections, particularly those caused by C. pneumoniae, including respiratory diseases, e.g. cough, sore throat, bronchitis, asthma. The polynucleotides, including DNA or RNA may be used in producing the encoded polypeptide in a recombinant host system, in the construction of vaccine vectors such as poxviruses, as vaccine agent, and in constructing attenuated Chlamydia strains that can over-express a polynucleotide or express it in a non-toxic mutated form. The polypeptides may also be used as diagnostic reagent for detecting the presence of anti-Chlamydia antibodies, and in the preparation of a medicament for treating or preventing Chlamydia infection. Dwg.0/4

FS CPI EPI AB; DCN FA CPI: B04-C01G; B04-E02F; B04-E03F; B04-E04; B04-E06; B04-E08; B04-F01; MC B04-F02; B04-G01; B04-G21; B04-G22; B11-C07; B11-C08; B11-C08E5; B12-K04A; B12-K04F; B14-A01A; B14-K01; B14-S11; D05-C07; D05-C11; D05-H07; D05-H08; D05-H09; D05-H12A; D05-H12B; D05-H12C; D05-H12D1; D05-H12D2; D05-H12D6; D05-H12E; D05-H14B2; D05-H17A1; D05-H17B; D05-H17B6; D05-H17C EPI: S03-E14H4 ANSWER 10 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN L54 2001-316102 [33] AN WPIX DNC C2001-097308 DNN N2001-227243 New Npt2cp (ADP/ATP translocase) polypeptides and nucleic acids encoding the polypeptides useful for treating, preventing or diagnosing Chlamydia infections, particularly infections caused by Chlamydia pneumoniae. B04 D16 S03 DC DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J IN (AVET) AVENTIS PASTEUR LTD

CYC 95 A1 20010329 (200133)\* EN 79 WO 2001021803 C12N015-31 PΙ RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ

NL OA PT SD SE SL SZ TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE  $\mathtt{SG} \ \mathtt{SI} \ \mathtt{SK} \ \mathtt{SL} \ \mathtt{TJ} \ \mathtt{TM} \ \mathtt{TR} \ \mathtt{TT} \ \mathtt{TZ} \ \mathtt{UA} \ \mathtt{UG} \ \mathtt{US} \ \mathtt{UZ} \ \mathtt{VN} \ \mathtt{YU} \ \mathtt{ZA} \ \mathtt{ZW}$ 

AU 2000073984 A 20010424 (200141) C12N015-31 A1 20020710 (200253) EN C12N015-31 EP 1220924

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

WO 2001021803 A1 WO 2000-CA1087 20000915; AU 2000073984 A AU 2000-73984 20000915; EP 1220924 A1 EP 2000-962124 20000915, WO 2000-CA1087 20000915 AU 2000073984 A Based on WO 2001021803; EP 1220924 A1 Based on WO FDT 2001021803

PRAI US 1999-154326P 19990917

ICM C12N015-31

ICS A61K031-711; A61K039-118; A61K039-40; C07K014-295; C07K016-12; C12N015-62; C12N015-85; G01N033-53

AB WO 200121803 A UPAB: 20010615

NOVELTY - A nucleic acid (I) encoding a polypeptide comprising:

(a) a fully defined sequence of 540 amino acids given in the specification:

(b) an immunogenic fragment comprising at least 12 consecutive amino acids from (a); or

(c) (a) or (b) which has been modified to improve its immunogenicity and which is at least 75% identical to (a) or (b), is new.

DETAILED DESCRIPTION - The nucleic acid has 1823 bp sequence given in the specification, or comprises at least 38 consecutive nucleotides of this sequence.

INDEPENDENT CLAIMS are also included for the following:

- (1) a nucleic acid comprising a sequence antisense to (I);
- (2) a nucleic acid encoding a fusion protein comprising a polypeptide encoded by (I) and an additional polypeptide;
- (3) vaccines comprising at least one first nucleic acid expressed as a polypeptide, a vaccine vector, and optionally a second nucleic acid encoding an additional polypeptide that enhances the immune response to

the polypeptide expressed by the first nucleic acid;

- (4) a unicellular host transformed with the nucleic acid;
- (5) a nucleic acid probe of 5-100 nucleotides or a primer of 10-40 nucleotides, which hybridizes under stringent conditions to an 1823-bp sequence, or to its homologue, complement, or antisense sequence;
  - (6) a polypeptide encoded by the nucleic acids;
- (7) a fusion polypeptide comprising a polypeptide of (6) and an additional polypeptide;
- (8) a method of producing a polypeptide of (6) by culturing a unicellular host of (4);
  - (9) an antibody against the polypeptide of (6);
- (10) pharmaceutical compositions comprising a polypeptide or an antibody:
- (11) a method of preventing or treating Chlamydia infection using the above nucleic acids, vaccines, pharmaceutical composition, polypeptides or antibodies;
- (12) a method of detecting Chlamydia infection by assaying a body fluid of a mammal with the above nucleic acids, polypeptides or antibody;
- (13) a diagnostic kit comprising instructions for use and the above nucleic acids, polypeptides or antibodies;
- (14) a method for identifying a polypeptide that induces immune response effective to prevent or lessen the severity of Chlamydia infection in a mammal previously immunized with the polypeptide by: (a) immunizing a mouse with the polypeptide; and (b) inoculating the immunized mouse with Chlamydia, where the polypeptide prevents or lessens the severity of Chlamydia infection in the immunized mouse compared to a non-immunized;
  - (15) expression plasmid pCABk663;
- (16) a nucleic acid having a 42 or 33 bp sequence given in the specification; and (17) Npt2cp (ADP/ATP translocase) from Chlamydia pneumoniae.

ACTIVITY - Antimicrobial.

MECHANISM OF ACTION - Vaccine. 7-9 week old male Balb/c mice were immunized intramuscularly plus intranasally with plasmid containing the coding sequence of Chlamydia pneumoniae Npt2cp (ADP/ATP translocase) or plasmid vector lacking an inserted chlamydial gene. Immunization was given at 0. 3 and 6 weeks, and at week 8, mice were inoculated with 5 multiply 105 IFU of C. pneumoniae strain AR39. 9 days post-challenge, lungs were taken and homogenized in SPG buffer. Dilutions of homogenate were assayed for the presence of infectious chlamydia by inoculation onto monolayers of susceptible cells. Cells were incubated for 3 days, and monolayers were fixed with formalin and methanol, and imunoperoxidase stained for the presence of chlamydial inclusions using convalescent sera from rabbits infected with C. pneumoniae. Mice immunized with pCABk663 had chlamydial lung titers less than 36,000 in 5 of 6 cases at day 9, while the range of values for the controls was 13,600-458,100 IFU/lung.

USE - The nucleic acids encoding the Chlamydia Npt2cp (ADP/ATP translocase) polypeptides are useful as a vaccine in preventing, treating or diagnosing Chlamydia infections, particularly those caused by C. pneumoniae, including respiratory diseases, e.g. cough, sore throat, bronchitis, asthma. The polynucleotides, including DNA or RNA may be used in producing the encoded polypeptide in a recombinant host system, in the construction of vaccine vectors such as poxviruses, as vaccine agent, and in constructing attenuated Chlamydia strains that can over-express a polynucleotide or express it in a non-toxic mutated form. The polypeptides may also be used as diagnostic reagent for detecting the presence of anti-Chlamydia antibodies, and in the preparation of a medicament for treating or preventing Chlamydia infection.

FS CPI EPI

FA AB: DCN

MC CPI: B04-C01G; B04-E02; B04-E02E; B04-E03E; B04-E06; B04-E08; B04-F01;

B04-G01; B04-G03; B04-L01; B11-C08E; B12-K04A4;

B14-A01A; D05-C03; D05-C07; D05-H07; D05-H09; D05-H11;

D05-H12A; D05-H12B; D05-H12C; D05-H12D2; D05-H12E; D05-H14; D05-H17A;

D05-H17B; D05-H17C

EPI: S03-E14H4

- L54 ANSWER 11 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
- AN 2001-257992 [26] WPIX
- DNN N2001-183971 DNC C2001-077792
- TI Novel Chlamydia pneumoniae lpdA protein and polynucleotides encoding them useful as component of vaccines for treating Chlamydia infections, and for detecting Chlamydia infection in the body fluid of a mammal.
- DC B04 D16 S03
- IN DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J

```
PA
     (AVET) AVENTIS PASTEUR LTD
CYC 94
                    A1 20010329 (200126) * EN
                                                78
     WO 2001021802
                                                       C12N015-31
PΙ
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TZ UG ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM
            DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC
            LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE
            SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
     AU 2000073983 A 20010424 (200141)
                                                       C12N015-31
ADT WO 2001021802 A1 WO 2000-CA1086 20000915; AU 2000073983 A AU 2000-73983
     20000915
     AU 2000073983 A Based on WO 2001021802
                          19990917
PRAI US 1999-154325P
    ICM C12N015-31
IC
     ICS
         A61K039-118; C07K014-295; C07K016-12; C07K019-00;
          C12N005-10; C12N015-62; C12N015-63; C12Q001-68; G01N033-53
     WO 200121802 A UPAB: 20010515
     NOVELTY - A polypeptide (I) which is (i) a polypeptide having fully
     defined Chlamydia pneumoniae lpdA protein sequence of 461 amino acids (S2)
     given in the specification, (ii) an immunogenic fragment of (S2)
     comprising 12 consecutive amino acids or (iii) polypeptide of (i) or (ii)
     which has been modified to improve its immunogenicity, and having 75% identity to amino acid sequence of (i) or (ii), is new.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the
          (1) a nucleic acid molecule (II) comprising a nucleic acid sequence
     which encodes (I);
          (2) a nucleic acid molecule (III) comprising a nucleic acid sequence
     which is antisense to (II);
          (3) a nucleic acid molecule (IV) comprising a nucleic acid sequence
     which encodes a fusion protein, comprising (I) encoded by (II) and an
     additional polypeptide;
          (4) a vaccine (V) comprising (I), (II) or (IV) and a vaccine vector,
     where each nucleic acid is expressed as a polypeptide. The vaccine
     optionally comprising a second nucleic acid encoding an additional
     polypeptide which enhances the immune response to the polypeptide
     expressed by the above mentioned nucleic acid;
          (5) a pharmaceutical composition comprising (II), (II) or (IV) and a
     carrier;
          (6) a unicellular host transformed with (II), (III) or (IV) which is
     operatively linked to one or more expression control sequences;
          (7) a nucleic acid probe (VI) of 5 to 100 nucleotides which
     hybridizes under stringent conditions to a fully defined C.pneumoniae lpdA
     gene sequence of 1586 nucleotides (S1) as given in the specification, its
     homolog or complementary or anti-sense sequence;
          (8) a primer of 10 to 40 nucleotides which hybridizes under stringent
     conditions to (S1), or to a homolog or complementary or anti-sense
     sequence of the nucleic acid molecule;
          (9) a polypeptide encoded by (II) or (IV);
          (10) a fusion polypeptide (VII) comprising (I) and an additional
    polypeptide;
          (11) preparation of (I);
          (12) an antibody (VIII) against (I);
          (13) a vaccine (IX) comprising (I) or (VII), and a carrier and
     optionally comprising a second polypeptide which enhances the immune
     response to (I);
          (14) a pharmaceutical composition comprising (I), (VII), (IX) or
     (VIII) and a carrier;
          (15) a diagnostic kit comprising instructions for use and (II),
     (III), (IV), (I), (VII) or (VIII);
          (16) identifying (I) or (VII) which induces an immune response
     effective to prevent or lessen the severity of Chlamydia infection in a
     mammal previously immunized with polypeptide involves immunizing a mouse
    with (I) or (VII) and inoculating the immunized mouse with Chlamydia;
          (17) expression plasmid pCABk892;
          (18) a nucleic acid molecule having a fully defined sequence of
     ataagaatgcggccgccaccatgacccaagaatttgattgtgttg (S3) or
     cggggtaccgtgacttaggagggaagtgtaaag (S4); and
          (19) lpdA protein from C.pneumoniae.

ACTIVITY - Antibacterial. The biological activity of (I) was tested
     in mice. Groups of 7 to 9 week old male Balb/c mice (6 to 10 per group)
     were immunized intramuscularly (i.m.) plus intranasally (i.n.) with
```

plasmid DNA containing the coding sequence of C.pneumoniae lpdA. At week 8, immunized mice were inoculated i.n. with 5 multiply 105 IFU of C.pneumoniae, strain AR39 in 100 mu l of SPG buffer to test their ability

to limit the growth of a sublethal C.pneumoniae challenge. Lungs were taken from the mice at day 9 post-challenge and immediately homogenized in SPG buffer (7.5% sucrose, 5 mM glutamate, 12.5 mM phosphate pH 7.5). Dilutions of the homogenate were assayed for the presence of infectious Chlamydia by inoculation onto monolayers of susceptible cells. The inoculum was centrifuged onto the cells, then the cells were incubated for three days at 35 deg. C in the presence of 1 mu g/ml cycloheximide. After incubation the monolayers were fixed with formalin and methanol then immunoperoxidase stained for the presence of chlamydial inclusions using convalescent sera from rabbits infected with C.pneumoniae and metal-enhanced DAB as a peroxidase substrate. Results showed that mice immunized i.n. and i.m. with pCABk892 had chlamydial lung titers less than 25,000 in 6 of 6 cases at day 9 whereas the range of values for control mice sham immunized with saline was 13,600-458,100 IFU/lung at day 9. MECHANISM OF ACTION - Vaccine; Gene therapy.
USE - (II), (III), (IV), (V), (VIII) or (IX) or the pharmaceutical

compositions as described above are useful for preventing or treating Chlamydia (C.trachomatis, C.psittaci, C.pneumonia or C.pecorum) infection. (I), (II), (III), (IV), (VII) or (VIII) is useful as diagnostic reagents for detecting Chlamydia infection which involves assaying a body fluid of a mammal to be tested for the above mentioned components. (II) is useful for producing (I) (claimed). The vaccine vectors, (I), (II), (VIII) are useful in the preparation of a medicament for preventing and/or treating Chlamydia infection. (VI) is useful in diagnostic tests as capture or detection probes. (VI) is thus useful as an agent for detecting and/or identifying presence of Chlamydia in the biological material. The primers derived from (II) are also useful for detecting and/or identifying Chlamydia in the biological material. (VIII) is also useful as a reagent for purifying (I) from a biological sample which involves carrying out antibody-based affinity chromatography with the biological sample. Dwg.0/4

```
CPI EPI
FS
```

FA

AB; DCN

CPI: B04-C01G; B04-E03F; B04-E04; B04-E05; B04-E06; B04-E08; B04-F01; MC B04-F10A; B04-G09; B04-N0300E; B04-N03A; B11-C07A; B11-C08E5; B12-K04A4; B12-K04F; B14-A01A; B14-S03; B14-S11B; D05-C11; D05-H09; D05-H11; D05-H12A; D05-H12C; D05-H12D1; D05-H12D2; D05-H12D5; D05-H12E; D05-H14A1; D05-H17A6; D05-H17C1; D05-H18 EPI: S03-E14H4

L54 ANSWER 12 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN

AN 2001-168447 [17] WPIX

DNC C2001-050284

Novel multivalent immunogenic composition for conferring protection against infection caused by Hameophilus influenzae and Moraxella catarrhalis comprises four antigens derived from each of the two microorganisms. DC

IN KLEIN, M H; LOOSMORE, S M; SASAKI, K; YANG, Y

PA (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD

CYC

ΡI WO 2001005424 A2 20010125 (200117) \* EN 58 A61K039-00 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW

> W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE

SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
AU 2000059586 A 20010205 (200128) A61K039-0 A61K039-00

A2 20020502 (200236) EN A61K039-116 R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

US 6391313 B1 20020521 (200239) A61K039-116 B 20031030 (200382) AU 767096 A61K039-00 A 20031219 (200404) A61K039-00

ADT WO 2001005424 A2 WO 2000-CA811 20000711; AU 2000059586 A AU 2000-59586 20000711; EP 1200122 A2 EP 2000-945494 20000711, WO 2000-CA811 20000711; US 6391313 B1 US 1999-353617 19990715; AU 767096 B AU 2000-59586 20000711; NZ 516819 A NZ 2000-516819 20000711, WO 2000-CA811 20000711

FDT AU 2000059586 A Based on WO 2001005424; EP 1200122 A2 Based on WO 2001005424; AU 767096 B Previous Publ. AU 2000059586, Based on WO 2001005424; NZ 516819 A Based on WO 2001005424

PRAI US 1999-353617 19990715 ICM A61K039-00; A61K039-116

ICS A61P031-04

Graser 10/030313 WO 200105424 A UPAB: 20010328 NOVELTY - A multivalent immunogenic composition (I) for conferring protection in a host against disease caused by both Hameophilus influenzae (HI) and Moraxella catarrhalis (MC) comprising four different antigens, of which at least one antigen is from HI and one antigen is from MC, is new. Additionally three of the antigens of (I) are adhesins, and one is from ACTIVITY - Auditory; antibacterial. MECHANISM OF ACTION - Vaccine. Groups of five BALB/C mice were immunized subcutaneously on days 1,29 and 43 with one of the mouse H91A Hin47 + rHMW + rHia + r200 kDa vaccines. Blood samples were taken on days 0, 14, 28, 42 and 56. Groups of five Hartley outbreed guinea pigs were immunized intramuscularly on days 1, 29 and 43 with the vaccine as described above. Blood samples were taken on days 0, 14, 28, 42 and 56. Anti-H91A Hin47, anti-rHMW, anti-rHia and anti-r200 kDa IgG antibody titers were determined by antigen specific enzyme linked immunosorbant assays (ELISAs). The results of the immunogenicity studies showed that the final bleed sera obtained from mice immunized with 0.3 mu g, or 3.0 mu g each of H91A Hin47 + rHMW + rHia with 0, 0.3, 1.0, 3.0 or 10.0 mu g of added r200 kDa, all had high antibody titers to H91A Hin47 component. The final bleed sera obtained from the mice immunized with 3.0 mu g each of H91A Hin47 + rHMW + rHia with 0, 0.3, 1.0, 3.0 or 10.0 mu g of added r200 kDa, all had high titer antibodies to the rHMW apparent enhancing or inhibiting effect on the anti-rHMW response with the addition of the r200 kDa component. Mice immunized with 0.3 mu g each of H91A Hin 47 + HMW + rHia with 0, 0.3, 1.0, 3.0 or 10.0 mu g of added r200 kDa, all had high titer antibodies to the rHia component. There was no apparent enhancing or inhibiting effect on the anti-rHia response with the addition of the r200 kDa component. The final bleed sera obtained from guinea pigs immunized with 25 mu g or 50 mu g each of H91A Hin47 + rHMW + rHia with 0, 25, 50 or 100 mu g of added r200 kDa, all had high titer antibodies to the H91A Hin47 component. Also final bleed sera obtained from guinea pigs immunized with 25 mu g or 50 mu g each of H91A Hin47 + rHMW + rHia with 0, 25, 50 or 100 mu g of added r200 kDa, all had titer antibodies to the rHMW component. There was no apparent enhancing or inhibiting effect on the anti-rHMW response upon the addition of the r200 kDa antigen. USE - (I) is useful for immunizing a host against infection caused by both HI and MC including otitis media (claimed). ADVANTAGE - The multivalent vaccine can confer protection against encapsulated and unencapsulated HI and MC diseased in a safe and efficient . manner. Dwg.0/14 FS CPI FA AB: DCN CPI: B04-B04C1; B14-A01; B14-A01A; B14-N02; MC B14-S11B; D05-C02; D05-H07; D05-H12F L54 ANSWER 13 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN 2000-687542 [67] WPTX AΝ DNC C2000-209327 Nucleic acids encoding a 76 kDa protein from Chlamydia pneumoniae, useful for vaccinating against Chlamydia infections. DC B04 D16 TN DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J (AVET) AVENTIS PASTEUR LTD; (DUNN-I) DUNN P; (MURD-I) MURDIN A PA D; (OOME-I) OOMEN R P; (WANG-I) WANG J CYC A2 20001109 (200067)\* EN PΙ WO 2000066739 90 C12N015-31 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2000043885 A 20001117 (200111) C12N015-31 EP 1177301 A2 20020206 (200218) EN C12N015-31 R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI JP 2002542827 W 20021217 (200312) C12N015-09

ADT WO 2000066739 A2 WO 2000-CA511 20000503; AU 2000043885 A AU 2000-43885 20000503; EP 1177301 A2 EP 2000-925004 20000503, WO 2000-CA511 20000503; JP 2002542827 W JP 2000-615762 20000503, WO 2000-CA511 20000503; US

A1 20030522 (200336)

NZ 515674 A 20031219 (200404) US 2004086525 A1 20040506 (200430)

US 2003095973

A61K039-40

C12N015-31 C07H021-04

```
2003095973 Al Provisional US 1999-132270P 19990503, Provisional US
     1999-141276P 19990630, US 2000-564479 20000503; NZ 515674 A NZ 2000-515674
     20000503, WO 2000-CA511 20000503; US 2004086525 A1 Provisional US
     1999-132270P 19990503, Provisional US 1999-141276P 19990630, Cont of US
     2000-564479 20000503, US 2003-608559 20030630
FDT AU 2000043885 A Based on WO 2000066739; EP 1177301 A2 Based on WO
     2000066739; JP 2002542827 W Based on WO 2000066739; NZ 515674 A Based on
     WO 2000066739
PRAI US 1999-141276P
                           19990630; US 1999-132270P
                                                            19990503;
     US 2000-564479
                           20000503; US 2003-608559
                                                            20030630
     ICM A61K039-40; C07H021-04; C12N015-09; C12N015-31
     ICS A61K031-70; A61K039-00; A61K039-02;
          A61K039-118; A61K039-38; A61K039-39;
          A61K039-395; A61K048-00; A61P009-10; A61P011-00; A61P011-02;
          A61P011-06; A61P031-04; C07K014-295; C07K016-12; C07K019-00; C12N001-15; C12N001-19; C12N001-21; C12N005-10; C12N015-11;
          C12N015-62; C12N015-85; C12P021-02; C12Q001-68; G01N033-53;
          G01N033-566; G01N033-569
```

WO 200066739 A UPAB: 20001223

NOVELTY - Nucleic acids (NAM1) encoding a 76 kDa protein (PEP1) from Chlamydia pneumoniae, is new. NAM1 and PEP1 have defined nucleotide and amino acid sequences ((I)-(VIII)) given in the specification.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are included for the following:

(1) a nucleic acid molecule (NAM1) comprising a nucleic acid sequence which encodes a polypeptide selected from:

(a) one of 3 defined amino acid sequences ((I)-(III)) given in the specification:

(b) a immunogenic fragment comprising at least 12 consecutive amino acids from (I)-(III); and

(c) the polypeptides of (a) and (b) which have been modified to improve their immunogenicity (the modified polypeptide is at least 75% identical in sequence to the corresponding polypeptides of (a) and (b);

(2) a nucleic acid molecule (I') comprising a sequence antisense to

(3) a nucleic acid molecule (NAM2) which encodes a fusion protein that comprises a polypeptide encoded by NAM1 and an additional polypeptide;

- (4) a vaccine (VAC1) comprising NAM1 and/or NAM2 and a vaccine vector (each nucleic acid molecule is expressed as a polypeptide and the vaccine may comprise additional nucleic acids encoding other polypeptides which enhance the immune response to the polypeptide expressed from NAM1 and/or
- (5) a unicellular host (UCH) transformed with NAM1 and NAM2 operatively linked to at least 1 expression control sequence;
- (6) a nucleic acid probe of 5-100 nucleotides which hybridizes under stringent conditions to (I) (or homolog, complementary or antisense sequences of (I));
- (7) a polypeptide (PEP1) encoded by NAM1 or NAM2; (8) a fusion polypeptide (PEP2) comprising PEP1 and an additional polypeptide;
  - (9) a method for producing PEP1 comprising culturing UCH;
- (10) an antibody (Ab) against PEP1 and/or PEP2; (11) a vaccine (VAC2) comprising PEP1 and/or PEP2 (the vaccine may comprise additional polypeptides which enhance the immune response to PEP1 and/or PEP2);
- (12) a diagnostic kit comprising NAM1, NAM2, PEP1, PEP2 and/or Ab and instructions for use;
- (13) a method for identifying polypeptides (either PEP1 or PEP) which induce an immune response that prevents or reduces the severity of Chlamydia infections in mammals previously immunized with the polypeptide, comprising:
  - (a) immunizing a mouse with the polypeptide; and
- (b) inoculating the immunized mouse with Chlamydia (the polypeptide which prevents or lessens the severity of the Chlamydia infection in the immunized mouse compared to a non-immunized control mouse is identified);
- (14) an expression plasmid selected from pCACPNM555a, pCAI555, pCAD76kDa; and
  - (15) an isolated 76 kDa protein (PEP3) from Chlamydia.

ACTIVITY - Bactericidal.

MECHANISM OF ACTION - Vaccine.

Mice immunized intranasally and intramuscularly with pCACPNM555a had Chlamydial lung titers less than 30000 IFU/lung in 5 of 6 cases at day 9 the range of values for control mice sham immunized with saline were 20800-323300 IFU/lung.

USE - NAM1, NAM2, PEP1, PEP2, VAC1, VAC2 and Ab may be used as

Graser 10/030313

Page 82

antigens for preventing and treating Chlamydia infection by vaccination. NAM1, NAM2, PEP1, PEP2 and Ab may also be used to detect Chlamydia infection in mammals by using them to assay body fluid (claimed) (e.g. in DNA hybridization assays and immunoassays). Dwg.0/9 FS CPI FA AB: DCN CPI: B04-B04C1; B04-C01; B04-E03F; B04-E04; B04-E05; B04-E06; MC B04-E08; B04-F0100E; B04-F10A; B04-G07; B04-N03A0E; B11-A; B11-C07A; B11-C08E; B11-C09; B12-K04A4; B12-K04E; B12-K04F; B14-A01A; B14-S11B; D05-A01A4; D05-A01B; D05-C12; D05-H04; D05-H07; D05-H08; D05-H09; D05-H11; D05-H12A; D05-H12C; D05-H12D; D05-H12E; D05-H14; D05-H17A5; D05-H17C; D05-H18 ANSWER 14 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN L54 2000-431500 [37] AN WPIX DNC C2000-131168 New immunogenic composition for conferring protection in a host against a disease caused by Haemophilus influenzae, comprises two different antigens of H. influenzae, where one of the antigens is an adhesin. DC KLEIN, M H; LOOSMORE, S M; YANG, Y IN (CONN-N) CONNAUGHT LAB LTD; (KLEI-I) KLEIN M H; (LOOS-I) LOOSMORE S M; PA (YANG-I) YANG Y; (AVET) AVENTIS PASTEUR LTD CYC 91 PΙ WO 2000035477 A2 20000622 (200037)\* EN 44 A61K039-102 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ TZ UG ZW W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2000015439 A 20000703 (200046) EP 1140158 A2 20011010 (200167) EN A61K039-102 R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI W 20021002 (200279) A61K039-102 JP 2002532433 55 A 20030829 (200365) NZ 512679 A61K039-102 <--AU 772882 B2 20040513 (200462) A61K039-102 ADT WO 2000035477 A2 WO 1999-CA1189 19991215; AU 2000015439 A AU 2000-15439 19991215; EP 1140158 A2 EP 1999-957822 19991215, WO 1999-CA1189 19991215; JP 2002532433 W WO 1999-CA1189 19991215, JP 2000-587796 19991215; NZ 512679 A NZ 1999-512679 19991215, WO 1999-CA1189 19991215; AU 772882 B2 AU 2000-15439 19991215 FDT AU 2000015439 A Based on WO 2000035477; EP 1140158 A2 Based on WO 2000035477; JP 2002532433 W Based on WO 2000035477; NZ 512679 A Based on WO 2000035477; AU 772882 B2 Previous Publ. AU 2000015439, Based on WO 2000035477 PRAI US 1998-210995 19981215 ICM A61K039-102 A61K039-05; A61K039-08; A61K039-10; A61K039-116; A61K039-13; A61K039-295; A61K039-39; A61P027-16; A61P031-04; C07K014-285 ICA C12N015-09 WO 200035477 A UPAB: 20000807 NOVELTY - A new immunogenic composition (I) for conferring protection in a host against a disease caused by Haemophilus influenzae, comprises at least two different antigens of Haemophilus influenzae, where at least one of the antigens is an adhesin. DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for a method of immunizing a host against disease caused by infection with Haemophilus influenzae, including otitis media, comprising administering to the host an immunoeffective amount of (I). ACTIVITY - Antibacterial. MECHANISM OF ACTION - Vaccine. H91A Hin47 is partially protective in the chinchilla model of otitis media, as described in the US Patent Number 5,506,139.

In this model, 1 to 2 year old chinchillas (Moulton Chinchilla Ranch, Rochester, Minnesota) were immunized intramuscularly (i.m.) on days 0, 14 and 28 with 30 micro g of H91A Hin47 adsorbed to alum, and challenged on day 44 with 50 to 350 colony forming units (cfu) of live organisms delivered into the middle ear space via the epitympanic bulla. Animals were monitored by tympanometry and middle ear fluid was collected 4 days post challenge, mixed with 200 micro 1 of BHI (undefined) medium and dilutions plated onto chocolate agar plates that were incubated for 24 hours at 37 deg. C. Convalescent animals or those mock-immunized with alum

alone, were used as controls. For the multi-component vaccine study, 50 micro q of H91A Hin47 was mixed with 50 micro q of recombinant HMW (rHMW) and chinchillas were immunized as described above. The results of the protection study indicate that there was still partial protection afforded in the intrabulla challenge model by the combination of H91A Hin47 and rHMW. USE - The two different antigens of H. influenzae, at least one of which is an adhesin, are useful in the manufacture of a vaccine for conferring protection against disease caused by infection with H. influenzae, including otitis media. (I) is used as a vaccine (all claimed) against diseases caused by H. influenzae infection. Dwg.0/12 FS CPI AB: DCN FA CPI: B04-N0300E; B14-A01A; B14-S11B; D05-H07 MC ANSWER 15 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN 2000-412339 [35] WPIX AN DNC C2000-125066 DNN N2000-308180 Nucleic acids encoding polypeptide antigens from Chlamydia useful for preventing, diagnosing and treating diseases such as community acquired pneumonia, bronchitis, sinusitis and asthmatic bronchitis, adult-onset asthma. nc B04 C06 D16 S03 IN MURDIN, A D; OOMEN, R P; WANG, J; JACOBSON, E L; JACOBSON, M K (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD; (JACO-I) PΑ JACOBSON E L; (JACO-I) JACOBSON M K CYC 91 A2 20000608 (200035)\* EN 173 PΙ WO 2000032794 C12N015-62 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ TZ UG ZW W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2000015405 A 20000619 (200044) A2 20010926 (200157) C12N015-62 EP 1135509 EN R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI US 2002123517 A1 20020905 (200260). A61K031-44 175 JP 2002531095 W 20020924 (200278) C12N015-00 NZ 512308 Α 20040130 (200414) C12N015-62 MX 2001005617 A1 20030401 (200415) A61K039-118 WO 2000032794 A2 WO 1999-CA1147 19991201; AU 2000015405 A AU 2000-15405 19991201; EP 1135509 A2 EP 1999-957785 19991201, WO 1999-CA1147 19991201; US 2002123517 Al Provisional US 1998-110428P 19981201, CIP of US 1999-452617 19991201, Div ex US 2000-549691 20000414, US 2002-113681 20020508; JP 2002531095 W WO 1999-CA1147 19991201, JP 2000-585425 19991201; NZ 512308 A NZ 1999-512308 19991201, WO 1999-CA1147 19991201; MX 2001005617 A1 WO 1999-CA1147 19991201, MX 2001-5617 20010601 FDT AU 2000015405 A Based on WO 2000032794; EP 1135509 A2 Based on WO 2000032794; US 2002123517 A1 CIP of US 6337065, Div ex US 6403619; JP 2002531095 W Based on WO 2000032794; NZ 512308 A Based on WO 2000032794; MX 2001005617 A1 Based on WO 2000032794 PRAI US 1998-110438P 19981201; US 1998-110339P 19981201; US 1998-110340P 19981201; US 1998-110427P 19981201; US 1998-110428P 19981201; US 1999-452617 19991201; US 2000-549691 20000414; US 2002-113681 20020508 ICM A61K031-44; A61K039-118; C12N015-00; C12N015-62 TC A61K031-7088; A61K039-385; A61K039-39; A61K039-395; A61K048-00; A61P031-00; C07K007-08; C07K014-295; C07K014-705; C07K016-12; C07K019-00; C12N001-15; C12N001-19; C12N001-21; C12N005-10; C12P021-02; C12Q001-68; G01N033-15; G01N033-50; G01N033-53; G01N033-569 WO 200032794 A UPAB: 20000725 NOVELTY - Nucleic acids (NAM1) encoding polypeptide (PEP1) antigens from Chlamydia, are new. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are included for the following: (1) a nucleic acid molecule (NAM1) comprising a sequence selected (a) nucleic acid sequences (N1)-(N10), which are defined sequences given in the specification; (b) a sequence encoding a polypeptide encoded by (N1) - (N10); (c) a sequence comprising at least 38 consecutive nucleotides from (N1) - (N10); and/or

Graser 10/030313

Page 84

(d) a sequence which encodes a polypeptide at least 75% identical in amino acid sequence to the polypeptides encoded by (N1)-(N10);

- (2) a nucleic acid molecule (NAM1') that comprises an antisense sequence to NAM1;
- (3) a nucleic acid molecule (NAM2) comprising a sequence encoding a fusion protein comprising the polypeptide encoded by NAM1 and an additional polypeptide;
- (4) a vaccine composition (VAC1) comprising a vaccine vector and NAM1 and/or NAM2 expressed as a polypeptide (the vaccine may comprise an additional polypeptide that enhances the immune response to the polypeptide expressed by NAM1);
- (5) a unicellular host (CELL1) transformed with either NAM1 and/or NAM2;
- (6) a nucleic acid probe of 5-100 nucleotides which hybridizes under stringent conditions to (N1)-(N10) (or homologs, complementary and/or antisense sequences of them);
- (7) a primer of 10-40 nucleotides which hybridizes under stringent conditions to (N1)-(N10) (or homologs, complementary and/or antisense sequences of them);
  - (8) a polypeptide (PEP1) encoded by NAM1 and/or NAM2;
- (9) a fusion peptide (PEP2) comprising PEP1 and an additional polypeptide;
- (10) a method (METH1) for producing PEP1 and/or PEP2 comprising culturing CELL1:
  - (11) an antibody (ANB1) against PEP1 and/or PEP2;
- (12) a method (METH2) for preventing and/or treating Chlamydia infections using NAM1 and/or NAM2, VAC1, PEP1 and/or PEP2 or ANB1;
- (13) a method (METH3) for detecting Chlamydia infection comprising assaying a body fluid of a mammal to be tested with either NAM1 and/or NAM2, PEP1 or ANB1;
- (14) a diagnostic kit comprising instructions for use and either NAM1 and/or NAM2, PEP1 or ANB1; and
- (15) a method (METH4) for identifying a PEP1 and/or PEP2 which induces an immune response that prevents or lessens the severity of a Chlamydia infection in a mammal previously immunized with the polypeptide, comprising:
  - (a) immunizing a mouse with the polypeptide; and
- (b) inoculating the immunized mouse with Chlamydia (the polypeptide which lessens or prevents the severity of the Chlamydia infection in the immunized mouse compared to a non-immunized control mouse is identified).

ACTIVITY - Bactericide.

No data given.

MECHANISM OF ACTION - Vaccine.

USE - The nucleic acids (and complementary sequences) may be used as diagnosite agents for detecting the presence of nucleic acids encoding Chlamydia antigens in samples according to standard methods, and therefore, for diagnosing Chlamydia infections. For example, they may be used as primers and probes for diagnostic polymerase chain reaction (PCR) assays. Antisense sequences may be used to down regulate expression of the proteins and may be used to treat infections. The nucleic acids may also be used to produce the protein antigens they encode according to standard recombinant DNA methodologies. The proteins may then be used as antigens for the production of antibodies (i.e. as vaccines) for preventing infection by Chlamydia. The antibodies may also be used as diagnostic reagents for detecting infections. Chlamydia is a pathogen implicated in the development of (for example) community acquired pneumonia, upper respiratory tract disease (especially bronchitis and sinusitis, asthmatic bronchitis, adult-onset asthma and acute exacerbations of asthma in adults.

```
Dwg.0/0
```

FS CPI EPI

FA AB; DCN

MC

AB; DCN

CPI: B04-B04C1; B04-C01; B04-E03F; B04-E04; B04-E05; B04-E06;

B04-E08; B04-F0100E; B04-F10A; B04-G07; B04-N03A0E; B04-P01B; B11-A;

B11-C07A4; B11-C08E; B11-C09; B12-K04A4; B12-K04F; B12-M05;

B14-A01A; B14-S11B; B14-S12; C04-B04C1;

C04-C01; C04-E03F; C04-E04; C04-E05; C04-E06; C04-E08; C04-F0100E;

C04-F10A; C04-G07; C04-N03A0E; C04-P01B; C11-A; C11-C07A4; C11-C08E;

C11-C09; C12-K04A4; C12-K04F; C12-M05; C14-A01A;

C14-S11B; C14-S12; D05-A01A4; D05-A01B; D05-C12; D05-H04;

D05-H07; D05-H08; D05-H09; D05-H11; D05-H12A; D05-H12C; D05-H12D;

D05-H12E; D05-H14; D05-H17A5; D05-H17C; D05-H18

EPI: S03-E14H4

L54 ANSWER 16 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN AN 2000-412330 [35] WPIX

Graser 10/030313 Page 85

```
DNN N2000-308175
                       DNC C2000-125057
     New polynucleotide encoding the Chlamydia 98 kiloDalton outer membrane
     protein, useful for preventing or treating Chlamydia infection.
DC
     B04 D16 S03
     DUNN, P; MURDIN, A D; OOMEN, R P; WANG, J
IN
PA
     (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD; (DUNN-I)
     DUNN P; (MURD-I) MURDIN A D; (OOME-I) OOMEN R P; (WANG-I) WANG J
CYC
    91
                    A1 20000608 (200035) * EN
PΙ
     WO 2000032784
                                               94
                                                       C12N015-31
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL
            OA PT SD SE SL SZ TZ UG ZW
         W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES
            FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS
            LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL
            TJ TM TR TT TZ UA UG UZ VN YU ZA ZW
     AU 2000037909
                    A 20000619 (200044)
     EP 1135501
                     A1 20010926 (200157) EN
                                                       C12N015-31
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
            RO SE SI
     US 2002094340
                    A1 20020718 (200254)
                                                       A61K039-118
                    W 20020924 (200278)
     JP 2002531093
                                                 93
                                                       C12N015-09
                    A1 20030821 (200356)
                                                       A61K039-02
     US 2003157124
    MX 2001005616
                    A1 20030401 (200415)
                                                       A61K039-118
                    A 20040625 (200445)
    NZ 529361
                                                       C12N015-63
ADT WO 2000032784 A1 WO 1999-CA1148 19991201; AU 2000037909 A AU 2000-37909
     19991201; EP 1135501 A1 EP 1999-957786 19991201, WO 1999-CA1148 19991201;
     US 2002094340 Al Provisional US 1998-113439P 19981223, Provisional US
     1999-132272P 19990503, US 1999-452380 19991201; JP 2002531093 W WO 1999-CA1148 19991201, JP 2000-585415 19991201; US 2003157124 A1
     Provisional US 1998-110439P 19981201, Provisional US 1999-132272P
     19990503, Cont of US 1999-452380 19991201, US 2002-324129 20021220; MX
     2001005616 A1 WO 1999-CA1148 19991201, MX 2001-5616 20010601; NZ 529361 A
     Div ex NZ 2000-512354 20000114, NZ 2000-529361 20000114
    AU 2000037909 A Based on WO 2000032784; EP 1135501 Al Based on WO
     2000032784; JP 2002531093 W Based on WO 2000032784; MX 2001005616 A1 Based
     on WO 2000032784; NZ 529361 A Div ex NZ 512354
                          19990503; US 1998-110439P
                                                          19981201:
PRAI US 1999-132272P
     US 1998-113439P
                          19981223; US 1999-452380
                                                          19991201;
     US 2002-324129
                          20021220
    ICM A61K039-02; A61K039-118; C12N015-09; C12N015-31;
IC
          C12N015-63
     ICS A61K031-711; A61K038-00; A61K039-39; A61K039-395;
          A61K048-00; A61P031-04; C07H021-04; C07K014-295; C07K014-705;
          C07K016-12; C07K019-00; C12N001-15; C12N001-19; C12N001-21;
          C12N005-10; C12N015-11; C12N015-62; C12N015-74; C12P021-02;
          C12Q001-68; G01N033-15; G01N033-50; G01N033-53; G01N033-569
    WO 200032784 A UPAB: 20000725
    NOVELTY - Isolated polynucleotide (N1) encoding the Chlamydia 98
     kiloDalton (kDa) outer membrane protein, known as CPN100640, is new.
          DETAILED DESCRIPTION - Isolated polynucleotide (N1) encoding the
     Chlamydia 98 kiloDalton (kDa) outer membrane protein, is new.
          N1 comprises a nucleic acid sequence selected from:
          (a) the 3050 (I) or 2808 (II) nucleotide sequence defined in the
     specification;
          (b) a sequence which encodes a polypeptide encoded by (I) or (II);
          (c) a sequence comprising at least 38 consecutive nucleotides from
     any one of the nucleic acid sequences of (a) and (b); and
          (d) a sequence which encodes a polypeptide which is at least 75%
     identical in amino acid sequence to the polypeptides encoded by (I) or
          INDEPENDENT CLAIMS are also included for the following:
          (1) a nucleic acid molecule (N2) comprising a nucleic acid sequence
     which encodes a polypeptide selected from:
          (a) the 936 (III) or 925 (IV) amino acid sequence defined in the
     specification;
          (b) an immunogenic fragment comprising at least 12 consecutive amino
     acids from a polypeptide of (a); and
```

(2) a nucleic acid molecule (N3) comprising a nucleic acid sequence which is anti-sense to the nucleic acid molecule of (1) or N1;

its immunogenicity, where the modified polypeptide is at least 75% identical in amino acid sequence to the corresponding polypeptide of (a)

or (b):

(c) a polypeptide of (a) or (b) which has been modified to improve

(3) a nucleic acid molecule (N4) comprising a nucleic acid sequence which encodes a fusion protein comprising a polypeptide encoded by N1 and an additional polypeptide;

- (4) a vaccine comprising at least one first nucleic acid of N1, N2 or N4 and a vaccine vector, where each first nucleic acid is expressed as a polypeptide, the vaccine optionally comprising a second nucleic acid encoding an additional polypeptide which enhances the immune response to the polypeptide expressed by the first nucleic acid;
- (5) a unicellular host transformed with a nucleic acid (N1, N2, N3 or N4) operatively linked to one or more expression control sequences;
- (6) a nucleic acid probe of 5 to 100 nucleotides which hybridizes under stringent conditions to N1 or N2, or to a homolog or complementary or anti-sense sequence of the nucleic acid molecule;
- (7) a primer of 10 to 40 nucleotides which hybridizes under stringent conditions to N1 or N2, or to a homolog or complementary or anti-sense sequence of the nucleic acid molecule;
  - (8) a polypeptide (P1) encoded by N1, N2 or N4;
- (9) a polypeptide (P2) comprising an amino acid sequence selected from:
  - (a) (III) or (IV);
- (b) an immunogenic fragment comprising at least 12 consecutive amino acids from a polypeptide of (a); and
- (c) a polypeptide of (a) or (b) which has been modified to improve its immunogenicity, where the modified polypeptide is at least 75% identical in amino acid sequence to the corresponding polypeptide of (a) or (b);
- (10) a fusion polypeptide (P3) comprising P1 or P2, and an additional polypeptide;
- (11) a method for producing P1 or P2, comprising culturing the unicellular host of (5);
  - (12) an antibody against P1, P2 or P3;
- (13) a vaccine comprising at least one first polypeptide of P1, P2 or P3, and optionally comprising a second polypeptide which enhances the immune response to the first polypeptide;
- (14) a method of detecting Chlamydia infection comprising the step of assaying a body fluid of a mammal to be tested, with a component selected from:
  - (a) N1, N2, N3 or N4;

  - (b) P1, P2 or P3; or(c) the antibody of (12);
- (15) a diagnostic kit comprising a component selected from those defined in the method of (14);
- (16) a method for identifying a polypeptide of P1, P2 or P3 which induces an immune response effective to prevent or lessen the severity of Chlamydia infection in a mammal previously immunized with polypeptide, comprising:
  - (a) immunizing a mouse with the polypeptide; and
- (b) inoculating the immunized mouse with Chlamydia, where the immunized mouse is compared with a non-immunized mouse control to identify the polypeptide; and
  - (17) expression plasmid pCAI640.

ACTIVITY - Antibacterial.

Groups of 7 to 9 week old male Balb/c mice (8 to 10 per group) were immunized intramuscularly (i.m.) plus intranasally (i.n.) with plasmid DNA containing the coding sequence of C. pneumoniae 98 kDa outer membrane protein gene. Saline or the plasmid vector lacking an inserted chlamydial gene was given to groups of control animals.

For i.m. immunization alternate left and right quadriceps were injected with 100 micro g of DNA in 50 micro l of phosphate buffered saline (PBS) on three occasions at 0, 3 and 6 weeks. For i.n. immunization, anaesthetized mice aspirated 50 micro 1 of PBS containing 50 micro g DNA on three occasions at 0, 3 and 6 weeks. At week 8, immunized mice were inoculated i. n. with 5 x 105 IFU (undefined) of C. pneumoniae, strain AR39 in 100 micro l of SPG (7.5 % sucrose, 5 mM glutamate, 12.5 mM phosphate pH 7.5) buffer to test their ability to limit the growth of a sublethal C. pneumoniae challenge.

Lungs were taken from mice at days 5 and 9 post-challenge and immediately homogenized in SPG buffer. Dilutions of the homogenate were assayed for the presence of infectious Chlamydia by inoculation onto monolayers of susceptible cells. The inoculum was centrifuged onto the cells at 3000 rotations per minute (rpm) for 1 hour, then the cells were incubated for three days at 35 deg. C in the presence of 1 micro g/ml cycloheximide. After incubation the monolayers were fixed with formalin and methanol then immunoperoxidase stained for the presence of chlamydial inclusions using convalescent sera from rabbits infected with C. pneumoniae and metal-enhanced DAB (undefined) as a peroxidase substrate.

Mice immunized i.n. and i.m. with pCAI640 had chlamydial lung titers less than 255,000 in 4 of 4 cases at day 5 and less than 423,200 in 4 of 4 cases at day 9 while the range of values for control mice immunized with

```
saline was 227,000-934,200 IFU/lung (mean 685,240) at day 5 and
     96,000-494,000 IFU/lung (mean 238,080) at day 9.
         MECHANISM OF ACTION - Vaccine.
          USE - The nucleic acids, proteins, antibodies and vaccines are useful
     for preventing or treating Chlamydia infection (claimed).
FS
    CPI EPI
FΑ
    AB; DCN
    CPI: B04-C01G; B04-E02F; B04-E03F; B04-E05; B04-E06; B04-E08; B04-F0100E; B04-F1100E; B04-G01; B04-G07; B04-N03A0E; B11-C08E;
MC
          B12-K04A4; B12-K04F; B14-A01A; B14-S11B; D05-H07;
          D05-H09; D05-H11; D05-H12A; D05-H12B2; D05-H12D1; D05-H12D2;
          D05-H12E; D05-H14A; D05-H17A6; D05-H17B6
    ANSWER 17 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
L54
ΔN
     2000-352521 [31]
                        WPTX
    C2000-107478
     Novel multivalent vaccine composition for use solely as a booster against
     in pre-sensitized subjects, to protect against e.g., diphtheria,
     poliomyelitis and tetanus.
    B04 D16
DC
IN
     CARTIER, J R; LAROCHE, P
PΑ
     (INMR) PASTEUR MERIEUX MSD; (AVET) AVENTIS PASTEUR; (AVET)
     AVENTIS PASTEUR MSD
CYC
                     A1 20000531 (200031)* FR
                                               11
                                                      A61K039-295
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
            RO SE SI
     WO 2000030678
                    A1 20000602 (200033) FR
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL
            OA PT SD SE SL SZ TZ UG ZW
         W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB
            GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU
            LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR
            TT UA UG US UZ VN YU ZA ZW
     AU 2000013901 A 20000613 (200043)
     EP 1131094
                     A1 20010912 (200155) FR
                                                      A61K039-295
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
            RO SE SI
     CZ 2001001860
                    A3 20020116 (200215)
                                                       A61K039-295
                                                                      <--
    ZA 2001003634
                    A 20020731 (200271)
                                                44
                                                       A61K000-00
    NZ 511933
                     A 20020927 (200272)
                                                       A61K039-295
     AU 759221
                     B 20030410 (200337)
                                                       A61K039-295
                                                                      <--
                    W 20030902 (200358)
    JP 2003525858
                                                31
                                                       A61K039-05
                                                                      <--
    EP 1131094
                    B1 20041103 (200475)
                                           FR
                                                       A61K039-295
         R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE
ADT EP 1004314 A1 EP 1998-122373 19981126; WO 2000030678 A1 WO 1999-FR2913
     19991125; AU 2000013901 A AU 2000-13901 19991125; EP 1131094 A1 EP
    1999-972544 19991125, WO 1999-FR2913 19991125; CZ 2001001860 A3 WO
     1999-FR2913 19991125, CZ 2001-1860 19991125; ZA 2001003634 A ZA 2001-3634
     20010504; NZ 511933 A NZ 1999-511933 19991125, WO 1999-FR2913 19991125; AU
     759221 B AU 2000-13901 19991125; JP 2003525858 W WO 1999-FR2913 19991125,
    JP 2000-583561 19991125; EP 1131094 B1 EP 1999-972544 19991125, WO
     1999-FR2913 19991125
FDT AU 2000013901 A Based on WO 2000030678; EP 1131094 Al Based on WO
     2000030678; CZ 2001001860 A3 Based on WO 2000030678; NZ 511933 A Based on
     WO 2000030678; AU 759221 B Previous Publ. AU 2000013901, Based on WO
     2000030678; JP 2003525858 W Based on WO 2000030678; EP 1131094 B1 Based on
     WO 2000030678
PRAI EP 1998-122373
                          19981126
    ICM A61K000-00; A61K039-05; A61K039-295
    ICS A61K039-08; A61K039-10; A61K039-13;
         A61K039-29; A61P025-00; A61P031-04
AB
         1004314 A UPAB: 20040511
    NOVELTY - Vaccine formulated for exclusive use as a repeat ('booster')
    vaccine in a population already having received a primary vaccine and/or
     sensitized against at least the poliovirus, Corynebacterium diphteriae
     and/or C. tetani, is new.
         DETAILED DESCRIPTION - Vaccine (I) formulated for exclusive use as a
    repeat ('booster') vaccine in a population already having received a
    primary vaccine and/or sensitized against at least the poliovirus,
    Corynebacterium diphteriae and/or C. tetani, comprises,
          (1) at least 1.2 mg/ml of aluminum salt;
          (2) antigens derived from at least the poliovirus; and
```

(3) a quantity of diphtheria anatoxin (DA) used as an antigen of C.

```
diphteriae comprising between 4-16 Flocculation units (Fu);
          ACTIVITY - Immunostimulant; anti-viral.
          MECHANISM OF ACTION - Vaccine. Three lots of 0.5 ml per vaccine were
     used in a study of 31 adults, the lots differed only in the quantity of
     diphtheria anatoxin, 2 (lot A), 5 (B) and 8 (C) Fu/ml, the adults were
     divided into 3 groups of 10, one for each lot. Each subject was injected
     into their deltoid muscle. No systemic reactions were reported for any of
     the lots, although localized reactions were noted over the first week in 8
     subjects in lot A, 6 subjects in lot B and 8 subjects in lot C. Symptoms
     included redness and swellings around the site of injection, although all
     symptoms disappeared without treatment and did not affect the quality of
     life of the subjects. No reactions were observed beyond the first week and no serious reactions were observed in any subjects. In addition the immune
     responses to the 5 antigens were excellent for all three lots, despite
     initially elevated titers due to the young age of the subjects and because of recent vaccinations, a booster effect was obtained for each antigen.
          ADVANTAGE - The vaccine (I) is specifically designed for use as a
     booster vaccine only, and as such avoids the reduced immunogenicity that
     occurs when administering reduced dosages of normal primary vaccines. The
     quantity of immunogenic diphtheria anatoxin used (10 Fu/ml) and allows
     optimal immunogenic protection while minimizing undesirable side-effects,
     such as allergic reactions to the antigens.
     Dwg.0/0
     CPT
     AB; DCN
     CPI: B04-B04M; B04-F11; B14-A01A; B14-A01A1; B14-A01B;
          B14-A02A4; B14-A02A5; B14-S11A; B14-S11B;
          D05-H07; D05-H08; D05-H14
     ANSWER 18 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
L54
     2000-350742 [30]
                         WPIX
     N2000-262745
                         DNC C2000-106768
DNN
     Isolated polynucleotide encoding a Chlamydia polypeptide useful to treat,
     diagnose and prevent disease caused by Chlamydia infection.
     DUNN, P L; MURDIN, A D; OOMEN, R P (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD; (DUNN-I)
     DUNN P L; (MURD-I) MURDIN A D; (OOME-I) OOMEN R P
     91
                     A1 20000504 (200030)* EN
     WO 2000024901
                                                   88
                                                         C12N015-31
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL
            OA PT SD SE SL SZ TZ UG ZW
         W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES
            FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS
            LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL
            TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
                      A 20000515 (200039)
     AU 9963593
                      A1 20010822 (200149)
                                             EN
                                                         C12N015-31
     EP 1124964
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
            RO SE SI
                      B1 20020611 (200244)
                                                         A61K039-118
     US 6403101
     US 2002091096
                      A1 20020711 (200248)
                                                         A61K048-00
     JP 2002528081
                      W 20020903 (200273)
                                                  104
                                                          C12N015-09
                     A1 20030701 (200366)
                                                         A61K039-02
     MX 2001004356
                                                         C12P021-06
     IIS 6642025
                      B2 20031104 (200374)
                                                         C12N015-31
                      A 20031219 (200404)
     NZ 511886
     AU 770905
                      B2 20040304 (200453)
                                                         C12N015-31
ADT WO 2000024901 A1 WO 1999-GB3565 19991028; AU 9963593 A AU 1999-63593
     19991028; EP 1124964 A1 EP 1999-951017 19991028, WO 1999-GB3565 19991028;
     US 6403101 B1 Provisional US 1998-106037P 19981028, Provisional US
     1999-154658P 19990920, US 1999-427501 19991026; US 2002091096 A1
     Provisional US 1998-106037P 19981028, Provisional US 1999-154658P
     19990920, Div ex US 1999-427501 19991026, US 2001-905119 20010713; JP
     2002528081 W WO 1999-GB3565 19991028, JP 2000-578453 19991028; MX 2001004356 A1 WO 1999-GB3565 19991028, MX 2001-4356 20010430; US 6642025
     B2 Provisional US 1998-106037P 19981028, Provisional US 1999-154658P
     19990920, Div ex US 1999-427501 19991026, US 2001-905119 20010713; NZ
     511886 A NZ 1999-511886 19991028, WO 1999-GB3565 19991028; AU 770905 B2 AU
     1999-63593 19991028
FDT AU 9963593 A Based on WO 2000024901; EP 1124964 A1 Based on WO 2000024901;
     JP 2002528081 W Based on WO 2000024901; MX 2001004356 Al Based on WO
     2000024901; US 6642025 B2 Div ex US 6403101; NZ 511886 A Based on WO
     2000024901; AU 770905 B2 Previous Publ. AU 9963593, Based on WO 2000024901
PRAI US 1999-427501
                           19991026; US 1998-106037P
                                                             19981028;
     US 1999-154658P
                           19990920; US 2001-905119
     ICM A61K039-02; A61K039-118; A61K048-00; C12N015-09;
```

FS

FA

MC

AN

TI

DC

IN PΑ

CYC

ΡI

C12N015-31; C12P021-06 ICS A61K038-00; A61K039-00; A61K039-39; A61P009-10; A61P011-00; A61P011-06; C07H021-04; C07K001-22; C07K014-295; C07K016-12; C12N001-15; C12N001-19; C12N001-21; C12N005-10; C12N015-62; C12P021-02; C12Q001-68; G01N033-53; G01N033-543; G01N033-566; G01N033-569 ICI C12N001-21; C12N015-09; C12P021-02; C12R001:01; C12R001:01; C12R001:01 WO 200024901 A UPAB: 20000624 NOVELTY - An isolated polynucleotide (N1) encoding a lorf2 protein of a strain of Chlamydia pneumoniae, is new.

DETAILED DESCRIPTION - An isolated polynucleotide (N1) has a nucleotide sequence which comprises:

(a) a defined nucleotide sequence (I) of 1550 base pairs or functional fragments of (I);

(b) a nucleotide sequence encoding a polypeptide with a sequence at least 75% homologous to (II) which has a defined protein sequence of 422 amino acids, or functional fragments; or

(c) a sequence capable of hybridizing under stringent conditions to a sequence comprising (I), or functional fragments.

INDEPENDENT CLAIMS are also included for the following:

- (1) an isolated polypeptide (P1) with a sequence at least 75% homologous to (II), or functional fragments of (II);
  - (2) a polypeptide P2 comprising P1 linked to a fusion polypeptide;
- (3) an expression cassette comprising N1 operably linked to a promoter;
  - (4) an expression vector comprising the expression cassette of (3);
  - (5) a host cell comprising the expression cassette of (3);
- (6) a method of producing a recombinant polypeptide with sequence (II) comprising culturing the host cell of (5) and recovering the polypeptide;
  - (7) a vaccine vector comprising the expression cassette of (3);
- (8) a pharmaceutical composition containing P1 and one or more known Chlamydia antigens;
- (9) a method for inducing an immune response in a mammal comprising administering the vaccine vector of (7) or a composition containing P1 to induce an immune response;
- (10) a polynucleotide probe reagent capable of detecting the presence of Chlamydia in biological material comprising a polynucleotide that hybridizes to N1 under stringent conditions;
- (11) a hybridization method for detecting the presence of Chlamydia in a sample comprising:
  - (a) obtaining polynucleotide from the sample;
- (b) hybridizing the obtained polynucleotide with the polynucleotide probe reagent of (10) under conditions allowing hybridization of the probe and the sample; and
  - (c) detecting any hybridization occurring;
- (12) an amplification method for detecting the presence of Chlamydia in a sample comprising:
- (a) obtaining polynucleotide from the sample;(b) amplifying the polynucleotide using one or more polynucleotide probe reagents of (10); and
  - (c) detecting the amplified polynucleotide;
- (13) a method for detecting the presence of Chlamydia in a sample comprising contacting the sample with a detecting reagent that binds to P1 in the sample and detecting the formed complex;
- (14) an affinity chromatography method for substantially purifying a polypeptide with sequence (II) comprises:
- (a) contacting a sample containing (II) with a detecting reagent that binds to the polypeptide to form a complex;
   (b) isolating the formed complex;

  - (c) dissociating the formed complex; and
  - (d) isolating the dissociated polypeptide; and
- (15) an antibody that immunospecifically binds Pl or a fragment or derivative of the antibody containing its binding domain.

ACTIVITY - Antibacterial.

MECHANISM OF ACTION - Vaccine.

Balb/c mice (7-9 weeks old) were immunized intramuscularly and intranasally with plasmid DNA containing the coding sequence of C. pneumoniae lorf2 gene. Control animals were given saline or the plasmid vector without the chlamydial gene. The intramuscular immunization comprised 100 micro g DNA in 50 micro l phosphate buffered saline (PBS) at 0, 3 and 6 weeks and the intranasal immunization comprised 50 micro g DNA in 50 micro l PBS at 0, 3 and 6 weeks. At week 8, immunized mice were inoculated intranasally with 5x105 inclusion forming units (IFU) of C. pneumoniae, strain AR39 in 100 micro 1 SPG (sucrose, glutamate, phosphate) buffer. Lungs were taken from the mice at day 9 post challenge and

Page 90

homogenized in SPG buffer, the homogenate was assayed for the presence of infectious chlamydia by inoculation onto monolayers of susceptible cells After incubation the monolayers were fixed and immunoperoxidase stained for the presence of chlamydial inclusions using convalescent sera from rabbits infected with C. pneumoniae and metal-enhanced DAB (not defined) as a peroxidase substrate. Mice immunized with the plasmid containing the lorf2 gene had an average chlamydial lung titer of 11050 IFU/lung compared to 111783 IFU/lung for the control mice immunized with saline. USE - The polynucleotides and polypeptides can be used as a vaccine for humans to treat or prevent disease caused by Chlamydia infection and P1, N1 or an antibody to P1 can be used to diagnose a Chlamydia infection. Dwg.0/4 CPI EPI FA AB; DCN MC CPI: B04-B04D5; B04-C01G; B04-E03F; B04-E05; B04-E08; B04-F0100E; B04-F10A; B04-G01; B04-G21; B04-G22; B04-N03A; B11-C07A; B11-C08E3; B11-C08E5; B12-K04A4; B12-K04F; B14-A01A; B14-S11B; D05-H04; D05-H07; D05-H09; D05-H11A; D05-H11B; D05-H12A; D05-H12D1; D05-H12E; D05-H14; D05-H17A6; D05-H18B EPI: S03-E14H4 L54 ANSWER 19 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN 2000-350688 [30] WPIX AN DNC C2000-106714 ΤI Chlamydia antigenes and the proteins they encode, useful for vaccinating against Chlamydia infections that affect the respiratory tract. IN MURDIN, A D; OOMEN, R P; WANG, J; DUNN, P (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD PA CYC 91 WO 2000024765 A2 20000504 (200030)\* EN 165 C07K014-00 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ TZ UG ZW W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2000012541 A 20000515 (200039) EP 1129202 A2 20010905 (200151) EN C12N015-62 R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI W 20020917 (200276) JP 2002530052 222 C12N015-09 MX 2001004291 A1 20030601 (200417) C07K014-00 WO 2000024765 A2 WO 1999-CA992 19991028; AU 2000012541 A AU 2000-12541 19991028; EP 1129202 A2 EP 1999-955602 19991028, WO 1999-CA992 19991028; JP 2002530052 W WO 1999-CA992 19991028, JP 2000-578335 19991028; MX 2001004291 A1 WO 1999-CA992 19991028, MX 2001-4291 20010427 FDT AU 2000012541 A Based on WO 2000024765; EP 1129202 A2 Based on WO 2000024765; JP 2002530052 W Based on WO 2000024765; MX 2001004291 A1 Based on WO 2000024765 19981102; US 1998-106034P PRAI US 1998-107035P 19981028: US 1998-106039P 19981028; US 1998-106042P 19981028; US .1998-106044P 19981028; US 1998-106072P 19981029: 19981029; US 1998-106074P 19981029; US 1998-106587P US 1998-106073P 19981029: US 1998-106087P 19981102: US 1998-106588P 19981102; US 1998-106589P 19981102; US 1998-107034P 19981102 ICM C07K014-00; C12N015-09; C12N015-62 ICS A61K038-00; A61K039-118; A61K039-395; A61K048-00; A61P031-04; C07K014-295; C07K016-12; C12N001-15; C12N001-19; C12N001-21; C12N005-10; C12Q001-68; G01N033-15; G01N033-50; G01N033-53; G01N033-569 WO 200024765 A UPAB: 20000624 AΒ NOVELTY - Nucleic acids (A) encoding Chlamydia antigenes and the proteins (B) they express, are new. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are included for the following:

- (1) a nucleic acid molecule (A) comprising a sequence encoding a polypeptide (B) selected from:
- (a) one of 19 defined amino acid sequences ((IIa) (IIs)) given in the specification;
- (b) an immunogenic fragment comprising at least 12 consecutive amino acids from (IIa) (IIs); and/or
- (c) a modified form of the polypeptide sequences (IIa) (IIs) which has been modified to improve its immunogenicity (the modified peptide is at least 75% identical to the corresponding amino acid sequence of (IIa) -

```
(IIb)):
          (2) a polypeptide (B) encoded by (A);
          (3) a nucleic acid encoding a fusion protein comprising a polypeptide
     encoded by (A) and an additional polypeptide;
          (4) a fusion protein comprising (B) and an additional polypeptide;
          (5) a vaccine (C) comprising (A) and a vaccine vector which express
     (B) (and optionally comprising a second nucleic acid (D) encoding an
     additional polypeptide (J) which enhances the immune response to (A)
     and/or (B);
          (6) a nucleic acid probe (E) (of 5 to 100 nucleotides) or primer (F)
     (of 10 to 40 nucleotides) which hybridizes under stringent conditions to
     (Ia) - (Iz) (or a homolog, complementary or antisense sequence of (Ia) -
     (Iz));
          (7) a unicellular host (G) transformed with (A);
          (8) a method for producing (B) comprising culturing (G);
          (9) a vaccine (H) comprising (B) (and optionally comprising an
     additional polypeptide (J));
          (10) an antibody (K) against (B);
          (11) a method for preventing or treating Chlamydia infection, using:
     (a) (A);
          (b) (C) and/or (H);
     (c) (B); and/or
     (d) (K);
          (12) a method for detecting Chlamydia infection comprising assaying a
     body fluid of a mammal with either:
     (b) (B); and/or
     (c) (K); and
          (13) a diagnostic kit comprising instructions for use and a component
     selected from:
     (a) (A);
     (b) (B); and/or
     (c) (K).
          ACTIVITY - Antiinflammatory; respiratory; antibacterial;
     anti-asthmatic; antiarteriosclerotic.
          No biological data given.
          MECHANISM OF ACTION - Vaccine.
          USE - The nucleic acids may be used for the recombinant production of
     the Chlamydia polypeptides (either in vivo or in vitro) according to
     standard recombinant DNA methodologies. The polypeptides may then be used
     to vaccinate against Chlamydia infections in mammals. Chlamydia, such as
     C. pneumoniae, are pathogens responsible for upper respiratory tract
     infections such as community acquired pneumonia, acute respiratory disease
     and bronchitis and may be implicated in atherosclerotic changes and
     asthma.
          The nucleic acids may also be used as probes for detecting the
     presence of Chlamydia nucleic acids in samples (and therefore diagnose
     infections) and the proteins may be used as antigens for the production of
     antibodies that may be used to detect Chlamydia proteins in samples (e.g.
     via enzyme linked immunosorbant assay (ELISA)).
     Dwg.0/0
     CPI
     AB: DCN
     CPI: B04-B03C; B04-B04C; B04-B04C1; B04-B04C7
          ; B04-B04M; B04-C01G; B04-E03F; B04-E04; B04-E05; B04-E06; B04-E08;
          B04-F01; B04-G07; B04-N03A0E; B11-A; B11-C07A4; B11-C08E1; B11-C08E3;
          B11-C08E5; B12-K04A4; B12-K04F; B14-A01A; B14-C03;
          B14-K01A; B14-S03; B14-S11B; D05-A01A4; D05-A01B; D05-C12;
          D05-H04; D05-H07; D05-H08; D05-H09; D05-H11; D05-H12A; D05-H12D1; D05-H12D2; D05-H12D5; D05-H12E; D05-H14; D05-H17A5; D05-H18B
     ANSWER 20 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
     2000-303789 [26]
                        WPIX
DNC C2000-092308
     Nucleic acid molecule for producing recombinant high molecular weight
     proteins of Haemophilus which are used as a vaccine to provide protection
     against Haemophilus induced diseases in humans.
     B04 D16
     KLEIN, M H; LOOSMORE, S M; YANG, Y
     (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD; (KLEI-I)
     KLEIN M H; (LOOS-I) LOOSMORE S M; (YANG-I) YANG Y
     90
                    A2 20000413 (200026) * EN 307
     WO 2000020609
                                                       C12N015-70
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL
            OA PT SD SE SL SZ TZ UG ZW
         W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES
```

FS

FA

MC

AN

TI

DC

IN

PA

CYC

PI

```
FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS
            LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ
            TM TR TT UA UG US UZ VN YU ZA ZW
                     A 20000426 (200036)
A2 20010725 (200143) EN
     AU 9960736
                                                        C12N015-70
     EP 1117807
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
            RO SE SI
                      B1 20020813 (200255)
     US 6432669
                                                        C12P021-06
     JP 2002532062
                     W 20021002 (200279)
                                                 307
                                                        C12N015-09
     US 2003133943
                     A1 20030717 (200348)
                                                        A61K039-02
     AU 765339
                     B 20030918 (200370)
                                                        C12N015-70
                     A 20040130 (200414)
     NZ 511360
                                                         C12N015-70
     JP 2004166710
                     A 20040617 (200440)
                                                  90
                                                        C12N015-09
ADT WO 2000020609 A2 WO 1999-CA938 19991007; AU 9960736 A AU 1999-60736
     19991007; EP 1117807 A2 EP 1999-947153 19991007, WO 1999-CA938 19991007; US 6432669 B1 CIP of US 1998-167568 19981007, US 1998-206942 19981208; JP
     2002532062 W WO 1999-CA938 19991007, JP 2000-574704 19991007; US
     2003133943 A1 Cont of US 1998-167568 19981007, US 2002-193764 20021211; AU
     765339 B AU 1999-60736 19991007; NZ 511360 A NZ 1999-511360 19991007, WO
     1999-CA938 19991007; JP 2004166710 A Div ex JP 2000-574704 19991007, JP
     2004-37346 20040213
FDT AU 9960736 A Based on WO 2000020609; EP 1117807 A2 Based on WO 2000020609;
     JP 2002532062 W Based on WO 2000020609; AU 765339 B Previous Publ. AU
     9960736, Based on WO 2000020609; NZ 511360 A Based on WO 2000020609
PRAI US 1998-206942
                           19981208; US 1998-167568
                                                           19981007:
                           20021211
     US 2002-193764
IC
     ICM A61K039-02; C12N015-09; C12N015-70; C12P021-06
     ICS A61K009-127; A61K009-14; A61K009-48; A61K031-70; A61K038-16;
          A61K039-102; A61P031-04; C07H021-04; C07K014-195;
          C07K014-285; C12N001-12; C12N001-21; C12N015-31; C12N015-74;
          C12P021-02
ICI C12P021-02; C12R001:93
     WO 200020609 A UPAB: 20000531
     NOVELTY - A nucleic acid molecule (I) comprising a promoter functional in
     Escherichia coli and operatively coupled to a modified operon of a
     non-typeable strain of Haemophilus comprising A, B and C genes, where the
     A gene only contains a nucleic acid sequence encoding a mature high
```

molecular weight protein (HMW) of the non-typeable strain of Haemophilus, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a vector adapted for transformation of a host comprising (I);
- (2) a strain of E. coli transformed by the vector of (1) expressing a protective HMW protein of a non-typeable strain of Haemophilus;
- (3) a recombinant protective HMW protein of a non-typeable strain of Haemophilus or immunogenic fragment or analog, produced by the transformed E. coli strain of (2);
- (4) a plasmid vector (II) for expression of a HMW protein of a non-typeable strain of Haemophilus comprising the T7 promoter, a cloning site for insertion of a nucleic acid molecule into the plasmid vector and portions B and C of the operon of a non-typeable Haemophilus strain;
- (5) an isolated and purified HMW 1 protein of a non-typeable strain of Haemophilus free from contamination by HMW 2 of the same strain of non-typeable Haemophilus;
- (6) an isolated and purified HMW 2 protein of a non-typeable strain of Haemophilus free from contamination by HMW 1 of the same strain of non-typeable Haemophilus;
- (7) an immunogenic composition comprising at least one immunogenically-active component which is (I), the recombinant protective HMW protein of (3) or the HMW proteins of (5) or (6) and a carrier;
- (8) a method for inducing protection against disease caused by Haemophilus comprising administering to a susceptible host the composition of (7); and
- (9) a method for producing a protective HMW protein of a non-typeable strain of Haemophilus comprising transforming E. coli with the vector of (1), growing the E. coli to express the encoded mature HMW protein and isolating and purifying the expressed HMW protein.

ACTIVITY - Antibacterial.

Groups of 8-9 chinchillas were immunized three times intramuscularly with 30 micro g of purified rHMWl or rHMW2, 2 x 109 colony forming units (cfu) of heat inactivated (56 deg. C for 10 minutes) H. influenzae (NTHi) whole cells in alum or alum, alone on days 0, 14 and 28. Serum samples and nasal wash samples were taken on day 42 for measurement of anti-HMWl or anti-HMW2 antibody titers by ELISAs (enzyme linked immunosorbent assays). On day 44, animals were lightly anesthetized using xylazine/ketamine hydrochloric acid by intramuscular injection. Intranasal inoculations were

performed by passive inhalation (0.1 ml per animal) of freshly cultured streptomycin-resistant NTHi strain 12 in BHI (not defined) medium supplemented with hemin and nicotinamide adenine dinucleotide (NAD) both at 2 micro g ml-1. Dose of bacterial challenge was 1 x 108 cfu per animal. Nasopharyngeal lavages were performed 4 days post inoculation on chinchillas. 67-88% of control animals immunized with alum only had culture positive nasal lavage fluids but 67-80% of animals immunized with the rHMW1 protein purified from constructs abc (pDS-1046-1-1), a/abc (pBK86-1-1) or abc/cer (pBK-76-1-1) were largely protected. Animals immunized with constructs that did not contain intact ABC genes were 70-90% infected. Similar results were achieved with rHMW2 protein. MECHANISM OF ACTION - Vaccine. USE - The nucleic acids and vectors are used for the production of recombinant H. influenzae HMW proteins which can be used as vaccines to mediate a humoral or cell-mediated immune response to provide protection against H. influenzae induced diseases in humans. The HMW proteins are also useful as antigens in immunoassays for detecting antibacterial, Haemophilus, HMW and/or peptide antibodies. The nucleotide sequences encoding the HMW proteins can be used to isolate and clone hmw genes from other non-typeable strains of Haemophilus in hybridization reactions. ADVANTAGE - Including the cer gene of E. coli enhances the level of expression of mature HMW protein by the vectors. Dwg.0/235 CPI AB: DCN CPI: B04-E03F; B04-E08; B04-F10A3E; B04-N03A0E; B14-A01A; B14-S11B; D05-H07; D05-H12A; D05-H12E; D05-H14A1; D05-H17A6 ANSWER 21 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN 2000-224701 [19] WPIX C2000-068762 Nucleic acid molecule encoding an inclusion membrane protein C of a strain of Chlamydia, useful as a vaccine for immunizing against Chlamydia B04 D16 DUNN, P L; MURDIN, A D; OOMEN, R P (CONN-N) CONNAUGHT LAB LTD; (DUNN-I) DUNN P L; (MURD-I) MURDIN A D; (OOME-I) OOMEN R P; (AVET) AVENTIS PASTEUR LTD 89 A1 20000302 (200019)\* EN 62 C12N015-31 WO 2000011181 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ UG ZW W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG US UZ VN YU ZA ZW A 20000314 (200031) AU 9953660 C12N015-31 Al 20010613 (200134) EN C12N015-31 EP 1105490 R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI B1 20030218 (200317) US 6521745 C07H021-04 US 6686339 B1 20040203 (200413) A61K048-00 A1 20041118 (200477) US 2004228874 C120001-68 WO 2000011181 A1 WO 1999-CA766 19990819; AU 9953660 A AU 1999-53660 19990819; EP 1105490 A1 EP 1999-939280 19990819, WO 1999-CA766 19990819; US 6521745 B1 Provisional US 1998-97199P 19980820, Provisional US 1999-132961P 19990507, US 1999-377399 19990820; US 6686339 B1 Provisional US 1998-97199P 19980820, Provisional US 1999-132961P 19990507, WO 1999-CA766 19990819, US 2001-763063 20010615; US 2004228874 A1 Provisional US 1998-97199P 19980820, Provisional US 1999-132961P 19990507, Div ex WO 1999-CA766 19990819, Div ex US 2001-763063 20010615, US 2004-756320 20040114 FDT AU 9953660 A Based on WO 2000011181; EP 1105490 A1 Based on WO 2000011181; US 6686339 B1 Based on WO 2000011181; US 2004228874 A1 Div ex US 6686339 PRAI US 1999-132961P 19990507; US 1998-97199P 19980820; US 1999-377399 19990820; US 2001-763063 20010615; US 2004-756320 20040114 ICM A61K048-00; C07H021-04; C12N015-31; C12Q001-68 ICS A61K035-66; A61K039-02; A61K039-118; C07K014-295; C07K016-12; C12N015-63 WO 200011181 A UPAB: 20000419 NOVELTY - An isolated and purified nucleic acid molecule (750 base pairs (bp)) (I) encoding an inclusion membrane protein C (203 amino acids) (II)

FS

FΑ

MC

L54 AN

DNC

TΙ

DC

IN

PA

CYC

PΙ

IC

AB

new.

of a strain of Chlamydia (both sequences given in the specification), is

Graser 10/030313 Page 94

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) an expression cassette containing (I);
- (2) an expression vector containing the expression cassette of (1);
  - (3) a vaccine vector containing (I).

ACTIVITY - Antibacterial.

Groups of 7 to 9 week old male Balb/c mice (8 to 10 per group) were immunized intramuscularly (i.m.) and intranasally (i.n.) with plasmid DNA containing (I). Saline was given to groups of control animals. For i.m. immunization, alternate left and right quadriceps were injected with 100 micro g of DNA in 50 micro 1 of phosphate buffered saline (PBS) on three occasions at 0, 3 and 6 weeks. For i.n. immunization, anaesthetized mice aspirated 50 micro 1 of PBS containing 50 micro g DNA on three occasions at 0, 3 and 6 weeks. At week 8, immunized mice were inoculated i.n. with 5 multiply 10 inclusion forming units (IFU) of Chlamydia pneumoniae, strain AR39 in 100 micro l of buffer to test their ability to limit the growth of a sublethal C. pneumoniae challenge. Lungs were taken from mice at days 5 and 9 post-challenge and homogenized. The homogenate was frozen at -70 deg. C until assay. Dilutions of the homogenate were assayed for the presence of infectious Chlamydia by inoculation into monolayers of susceptible cells. The inoculum was centrifuged onto the cells at 300 revolutions per minute (rpm) for 1 hour, the cells were then incubated for three days at 35 deg. C in the presence of 1 micro g/ml cycloheximide. After incubation, the monolayers were fixed with formalin and methanol then immunoperoxidase stained for the presence of chlamydial inclusions using convalescent sera from rabbits infected with C. pneumoniae. Mice immunized i.n. and i.m. with pCAII15 had chlamydial lung titers less than 262500 in 4 of the 4 cases at day 5 and less than 250000 in 4 of the 4 cases at day 9. In contrast, mice sham immunized with saline had 202400 to 886800 IFU/lung (mean 429800) at day 5 and 78400-284600 IFU/lung (mean 157080) at day 9.

MECHANISM OF ACTION - Vaccine.

USE - (I) is useful for protecting against Chlamydia infection.

Dwg.0/4 ĖS CPI

FΑ AB: DCN

MC

CPI: B04-B03C; B04-C01G; B04-E03; B04-E04; B04-E05; B04-E08;

B04-G01; B04-N03A; B11-C08D2; B11-C08E; B12-K04A4;

B14-A01A; B14-S11B; D05-H07; D05-H09; D05-H11;

D05-H12A; D05-H12D1; D05-H12D5; D05-H12E; D05-H17A

ANSWER 22 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN

AN 2000-224700 [19] WPIX

DNC C2000-068761

New nucleic acid encoding POMP91A protein from a strain of Chlamydia TI useful for preventing, treating and diagnosing Chlamydia infection.

DC

IN

DUNN, P L; MURDIN, A D; OOMEN, R P (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD PΑ

CYC 89

ΡI WO 2000011180 A1 20000302 (200019)\* EN 98 C12N015-31

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL

OA PT SD SE SL SZ UG ZW

W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ

TM TR TT UA UG US UZ VN YU ZA ZW

A 20000314 (200031) A1 20010613 (200134) EN AII 9953659 C12N015-31 EP 1105489 C12N015-31

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT

RO SE SI B1 20040217 (200413) US 6693087 A61K039-395

ADT WO 2000011180 A1 WO 1999-CA765 19990819; AU 9953659 A AU 1999-53659 19990819; EP 1105489 A1 EP 1999-939279 19990819, WO 1999-CA765 19990819;

US 6693087 B1 Provisional US 1998-97198P 19980820, US 1999-377850 19990820 AU 9953659 A Based on WO 2000011180; EP 1105489 A1 Based on WO 2000011180

PRAI US 1998-97198P 19980820; US 1999-377850 19990820

ICM A61K039-395; C12N015-31

ICS A61K031-70; A61K048-00; C07H021-04; C07K014-295

WO 200011180 A UPAB: 20000419

NOVELTY - Isolated and purified nucleic acid molecule (I) encoding a POMP91A protein or polypeptide fragment of POMP91A from a strain of Chlamydia, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

Graser 10/030313 Page 95

(1) an expression cassette containing (I) under the control of elements required for expression of (I);

- (2) an expression vector containing the expression cassette of (1);
- (3) a vaccine vector comprising (I) under the control of elements required for expression of (I); and
- (4) an antibody that specifically binds to a polypeptide with a protein sequence (II) of 947 amino acids or a polypeptide fragment containing the binding domain of (II).

ACTIVITY - Antibacterial.
MECHANISM OF ACTION - Vaccine.

Male Balb/c mice (7-9 weeks old) were immunized intramuscularly (100 micro g DNA in 50 micro 1 phosphate buffered saline) and intranasally (50 micro g DNA in 50 micro l phosphate buffered saline) with plasmid DNA pCAI327 containing the coding sequence of C. pneumonia POMP91A at 0, 3 and 6 week intervals. Control animals were given saline and plasmids pCAI116 and pCAI178 which express non-protective chlamydial antigens. After 8 weeks the immunized mice were inoculated intranasally with 5 x 105 IFU of C. pneumonia strain AR39 in 100 micro l SPG (sucrose, phosphate, qlutamate) buffer. The lungs were taken from the mice at days 5 and 9 post challenge and homogenized in SPG buffer (7.5% sucrose, 5 mM glutamate, 12.5 mM phosphate pH 7.5). Dilutions of the homogenate were assayed for the presence of infectious Chlamydia by inoculation onto monolayers of susceptible cells. The cells were incubated for 3 days at 35 deg. C in the presence of 1 micro g/ml cycloheximide and then fixed with formalin and methanol then immunoperoxidase stained using convalescent sera from rabbits infected with C. pneumonia and metal enhanced DAB (not defined) as peroxidase substrate. Mice immunized with pCAI327 had chlamydial lung titers less than 21500 in 5 of 6 cases at day 9 but for saline immunized mice the average titer was 49069 IFU/lung.

USE - (I) is used to prevent, treat and diagnose Chlamydia infection. Vaccine vectors containing (I) are used to induce an immune response against Chlamydia. (I) or a monoclonal antibody specific to POMP91A can be used to diagnose the presence of Chlamydia in a biological sample. Dwg.0/41

```
FS CPI
FA AB; DCN
```

MC CPI: B04-C01G; B04-E03F; B04-E08; B04-F0100E; B04-F10A; B04-G01; B04-N03A; B11-C07A; B11-C08E5; B12-K04A4; B12-K04F; B14-A01A; B14-S11B; D05-H09; D05-H11A; D05-H12A; D05-H12E; D05-H14; D05-H17A6

L54 ANSWER 23 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN

AN 2000-096387 [08] WPIX

CR 1995-194089 [25]; 1997-052329 [05]; 1998-100410 [09]; 1999-404437 [34]; 1999-404459 [34]; 1999-404487 [34]; 2000-181144 [16]

DNC C2000-027984

TI Antibodies specific for transferrin receptor proteins of Haemophilus influenzae, useful for treating otitis media, epiglottitis, pneumonia and tracheobronchitis.

DC B04 D16

IN CHONG, P; GRAY-OWEN, S; HARKNESS, R; KLEIN, M; LOOSMORE,
S; MURDIN, A; SCHRYVERS, A; YANG, Y

PA (CONN-N) CONNAUGHT LAB LTD

CYC 1

PI US 6008326 A 19991228 (200008)\* 252 C07K016-12 ADT US 6008326 A CIP of US 1993-148968 19931108, CIP of US 1993-175116

19931229, US 1995-474671 19950607, Cont of US 1995-337483 19951108 PRAI US 1995-337483 19951108; US 1993-148968 19931108; US 1993-175116 19931229; US 1995-474671 19950607

IC ICM C07K016-12

ICS C07K016-28

AB US 6008326 A UPAB: 20000925

NOVELTY - An isolated and purified antibody (or monospecific antiserum) specific for a single transferrin receptor protein (or immunogenic fragment) of a strain of Haemophilus influenzae, is new.

ACTIVITY - Antibacterial; antiinflammatory; auditory; respiratory. No relevant biological data given.

MECHANISM OF ACTION - Vaccine (antibody inhibition of bacterial growth and replication).

USE - The antibodies may be used for preventing and treating infections and disorders caused by H. influenzae, these include bacterial meningitis, otitis media, epiglottitis, pneumonia and tracheobronchitis. The antibodies may also be used detect the presence of H. influenzae proteins in samples according to standard methodologies (e.g. enzyme linked immunosorbant assay (ELISA)) and hence diagnose infections.

ADVANTAGE - The use of antibodies to treat bacterial infections

```
avoids the risk of antibiotic resistant bacterial strains developing. In
     the treatment of otitis media, the use of antibodies avoids the need for
     extensive and costly surgery (e.g. tonsillectomies, adenoidectomies and
     the insertion of tympanostomy tubes) to rectify hearing problems.
     Dwg.0/30
FS
     CPI
     AB; DCN
FA
MC
     CPI: B04-B04C1; B04-B04C7; B04-C01; B04-E03D;
          B04-F0100E; B04-G04; B04-G07; B04-G21; B04-K01T; B04-N0300E; B11-A;
          B11-C07A1; B11-C07A4; B11-C08E; B11-C09; B12-K04A4; B12-M05;
          B14-A01A; B14-C03; B14-K01; B14-N02; B14-N05;
          B14-S11B; D05-A01A4; D05-A01B; D05-C12; D05-H04; D05-H07;
          D05-H09; D05-H11A; D05-H12A; D05-H14; D05-H17A4; D05-H17A5
L54 ANSWER 24 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
     1999-620376 [53]
                        WPIX
AN
CR
     1997-457533 [42]
DNC
    C1999-181129
     Nucleic acid encoding transferrin binding protein 2 of Moraxella
     catarrhalis, useful for diagnostics, immunization and recombinant protein
     production.
DC
     B04 D16
     DU, R; HARKNESS, R E; KLEIN, M H; LOOSMORE, S M;
IN
     MYERS, L E; SCHRYVERS, A B; YANG, Y
PΑ
     (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD
CYC
    83
     WO 9952947
                     A2 19991021 (199953) * EN 113
                                                        C07K014-79
PΙ
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL
            OA PT SD SE SL SZ UG ZW
         W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GE
            GH GM HR HU ID IL IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG
            MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG
            US UZ VN YU ZW
                     A 19991101 (200013)
     AU 9931350
     EP 1071715
                     A2 20010131 (200108)
                                            EN
                                                        C07K014-79
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
            RO SE SI
                     A 20011016 (200170)
W 20020416 (200242)
     BR 9909576
                                                        C07K014-79
                                                        C07K014-705
     JP 2002511490
                                                 122
     US 6440701
                     B1 20020827 (200259)
                                                        C12N015-31
                     B 20030529 (200346)
A 20030725 (200357)
     AU 761008
                                                        C07K014-79
     NZ 507978
                                                        C07K014-79
ADT WO 9952947 A2 WO 1999-CA307 19990412; AU 9931350 A AU 1999-31350 19990412;
     EP 1071715 A2 EP 1999-913049 19990412, WO 1999-CA307 19990412; BR 9909576
     A BR 1999-9576 19990412, WO 1999-CA307 19990412; JP 2002511490 W WO
     1999-CA307 19990412, JP 2000-543503 19990412; US 6440701 B1 CIP of US 1996-613009 19960308, CIP of US 1997-778570 19970103, CIP of WO 1997-CA163
     19970307, US 1998-59584 19980414; AU 761008 B AU 1999-31350 19990412; NZ
     507978 A NZ 1999-507978 19990412, WO 1999-CA307 19990412
FDT AU 9931350 A Based on WO 9952947; EP 1071715 A2 Based on WO 9952947; BR
     9909576 A Based on WO 9952947; JP 2002511490 W Based on WO 9952947; AU
     761008 B Previous Publ. AU 9931350, Based on WO 9952947; NZ 507978 A Based
     on WO 9952947
PRAI US 1998-59584
                          19980414; US 1996-613009
                                                           19960308:
     US 1997-778570
                          19970103; WO 1997-CA163
                                                           19970307
     ICM C07K014-705; C07K014-79; C12N015-31
IC
     ICS A61K039-02; A61K048-00; C07K014-22; C12N001-15; C12N001-19;
          C12N001-21; C12N005-10; C12N015-09; C12N015-63; C12P021-02;
          C12Q001-68
ICI C07K014-705; C12N015-09; C12R001:01
         9952947 A UPAB: 20030906
     NOVELTY - Purified, isolated nucleic acid (I) encoding a transferrin
     binding protein (Tbp2) (II) from Moraxella catarrhalis strains
     M35, 3 or LES1, is new.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the
     following:
          (a) vectors containing (I);
          (b) transformed host cells containing the vector of (a);
          (c) recombinant production of (II);
          (d) recombinant (II) produced this way;
          (e) an immunogenic composition containing (I) or recombinant (II)
     plus a carrier;
          (f) a method for detecting Moraxella nucleic acid that
     encodes transferrin receptor protein by the formation of a hybrid with
          (g) diagnostic kits for the method of (f).
```

```
ACTIVITY - Antibacterial; cytostatic; auditory.
          MECHANISM OF ACTION - Tbp binding blocker.
          (I) and (II) generate an immune response that includes anti-Tbp
     antibodies and opsonizing and/or bactericidal antibodies. By blocking
    binding to Tbp, the antibodies stop the bacterium from acquiring essential
         USE - (I) is used to produce recombinant (II); for identification or
     diagnosis of Moraxella, or for cloning related species, using
     hybridization assays; and for genetic immunization against
     Moraxella infections, e.g. otitis media. (II) are useful as
     antigens, either in vaccines (including components of conjugate vaccines
     that contain antigens from other bacteria or from tumors, in which case
     they elicit production of antitumor antibodies that may be coupled to
     chemotherapeutic agents or biologically active agents) or to raise
     antibodies (for use as diagnostic reagents and for treating
    Moraxella infections), also for detecting Moraxella
     antibodies.
     Dwg.0/9
    CPI
    AB; DCN
FA
     CPI: B04-C01G; B04-E02F; B04-E05; B04-E08; B04-F0100E; B04-G01;
MC
          B04-K01T; B11-C07A; B11-C07A1; B11-C08E5; B12-K04A; B12-K04E;
          B12-K04F; B14-A01; B14-H01; B14-N02; D05-H04; D05-H07; D05-H09;
         D05-H11; D05-H12A; D05-H12E; D05-H14; D05-H17A4; D05-H17A6; D05-H18B
L54 ANSWER 25 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
    1997-373222 [35] WPIX
DNN N1997-309929
                       DNC C1997-120314
    Lactoferrin receptor protein isolated from bacterial pathogen - used as,
ΤI
     e.g. vaccine, carrier for antigens and immunogens and diagnostic agents.
DC
    A96 B04 D16 S03
IN
    BONNAH, R A; SCHRYVERS, A B
     (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD
PA
CYC 2
                    A 19970503 (199735) *
A 20000411 (200025)
PΤ
    CA 2162193
                                                50
                                                      C12P021-08
                                                      A61K039-095
     US 6048539
    US 6211343
                    B1 20010403 (200120)
                                                      C08H001-00
                    B1 20020205 (200211)
                                                      A61K039-02
     US 6344200
                                                                     <--
     US 6348198
                    B1 20020219 (200221)
                                                      A61K039-095
ADT CA 2162193 A CA 1995-2162193 19951106; US 6048539 A US 1995-552232
     19951102; US 6211343 B1 Div ex US 1995-552232 19951102, US 1999-370869
     19990810; US 6344200 B1 Div ex US 1995-552232 19951102, US 1999-371126
     19990810; US 6348198 B1 Div ex US 1995-552232 19951102, US 1999-371127
FDT US 6211343 B1 Div ex US 6048539; US 6344200 B1 Div ex US 6048539; US
     6348198 B1 Div ex US 6048539
PRAI US 1995-552232
                         19951102; US 1999-370869
                                                         19990810:
    US 1999-371126
                         19990810; US 1999-371127
                                                         19990810
    ICM A61K039-02; A61K039-095; C08H001-00; C12P021-08
    ICS C07K001-36; C07K014-22; C07K016-12; G01N033-566; G01N033-569
         2162193 A UPAB: 19970828
AB
    Lactoferrin receptor protein (I) is isolated and purified from a bacterial
    protein and has molecular weight (MW) 70-90 kDa as determined by SDS-PAGE.
         USE - The immunogenic composition can be used as a vaccine to a
    bacterial pathogen selected from Neisseria meningitides, N-gonorrhoeae,
    Moraxella catarrhalis, M. movis and M. lacunata (all claimed).
          The proteins can be used in the diagnosis of and vaccination against
     diseases caused by bacterial pathogens that produce lactoferrin receptor
    proteins or proteins capable of raising antibodies reactive with
     lactoferrin receptor proteins. The proteins can be used as antigens,
     immunogenic preparations including vaccines, carriers for other antigens
     and immunogens and the generation of diagnostic reagents.
          The bacterial pathogen may also be selected from Haemophilus
     influenzae, Streptococcus pneumoniae, Escherichia coli, Salmonella typhi,
     Streptococcus mutans, Cryptococcus neoformans, Klebsiella sp.,
     Staphylococcus aureus and Pseudomonas aeruginosa.
          (I; Lbp2) may also be used to induce immunity toward abnormal
     polysaccharides of tumour cells and to produce antitumour antibodies that
     can be conjugated to chemotherapeutic and bioactive agents.
    Dwg.0/4
    CPI EPI
FA
    AB
    CPI: A12-V01; A12-V03C2; B04-G01; B04-N03; B11-C07A; B12-K04A;
MC
         B14-S11B; D05-H07; D05-H09; D05-H13
     EPI: S03-E14H4
```

Page 98

```
L54 ANSWER 26 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
                         WPIX
AN
     1996-342057 [34]
     N1996-287919
                         DNC C1996-108616
DNN
TI
     New immunogenic conjugate molecules - comprise a capsular polysaccharide
     of Streptococcus linked to an outer membrane protein of Haemophilus.
DC
     FAHIM, R E F; GISONNI, L; KANDIL, A; KLEIN, M H; YANG, Y
IN
     ; FAHIM, R E
PA
     (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD
CYC
     71
                      A2 19960718 (199634) * EN 64
                                                          A61K039-09
     WO 9621465
PΙ
        RW: AT BE CH DE DK EA ES FR GB GR IE IT KE LS LU MC MW NL OA PT SD SE
             SZ UG
          W: AL AM AT AU AZ BB BG BR BY CA CH CN CZ DE DK EE ES FI GB GE HU IS
             JP KE KG KP KR KZ LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT
             RO RU SD SE SG SI SK TJ TM TR TT UA UG US UZ VN
     AU 9643254
                      A 19960731 (199645)
                                                          A61K039-09
     WO 9621465
                      A3 19961010 (199648)
                                                          A61K039-09
     US 5681570
                      A 19971028 (199749)
                                                   21
                                                          A61K039-385
                                                                          <--
                      A1 19971112 (199750) EN
     EP 805691
                                                          A61K039-09
                                                                          <--
         R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE
     US 6177085
                      B1 20010123 (200107)
                                                          A61K039-09
     US 6329512
                      B1 20011211 (200204)
                                                          A23J001-00
TOA
     WO 9621465 A2 WO 1996-CA7 19960105; AU 9643254 A AU 1996-43254 19960105;
     WO 9621465 A3 WO 1996-CA7 19960105; US 5681570 A US 1995-371965 19950112;
     EP 805691 A1 EP 1996-900066 19960105, WO 1996-CA7 19960105; US 6177085 B1
     Cont of US 1995-371965 19950112, US 1995-467884 19950606; US 6329512 B1
Cont of US 1995-371965 19950112, US 1995-467883 19950606
FDT AU 9643254 A Based on WO 9621465; EP 805691 A1 Based on WO 9621465; US
     6177085 B1 Cont of US 5681570; US 6329512 B1 Cont of US 5681570
PRAI US 1995-371965
                           19950112; US 1995-467884
                                                             19950606:
     US 1995-467883
                           19950606
     1.Jnl.Ref; EP 172107; EP 338265; EP 389925; EP 497524; EP 497525; US 4673574; US 5098997; WO 9106652
REP
     ICM A23J001-00; A61K039-09; A61K039-385
     ICS A61K039-02; A61K039-102; A61K039-116; C07K014-285; C07K017-10; G01N033-569
AB
     WO
          9621465 A UPAB: 19960829
     The following are claimed: (A) an immunogenic conjugate molecule (ICM)
     comprising at least a portion of a capsular polysaccharide (CP) of a
     Streptococcus strain linked to at least a portion of an outer membrane
     protein (OMP) of a Haemophilus strain, which are selected to provide in
     the ICM an enhanced immune response to the CP; (B) a diagnostic kit for
     determining the presence of antibodies in a sample specifically reactive
     with a CP of a Streptococcus strain or with an OMP of a Haemophilus
     strain; (C) a diagnostic kit for detecting the presence of a CP of a
     Streptococcus strain in a sample; (D) a diagnostic kit for detecting the
     presence of an OMP of a Haemophilus strain in a sample; (E) a process for
     individually isolating P1, P2 and P6 OMPs from a Haemophilus strain; and (F) an ICM comprising a P6 OMP of a Haemophilus strain linked to at least
     a portion of a CP of an encapsulated pathogen to provide in the ICM an
     enhanced immune response to the CP.
          The ICM is pref. used with an adjuvant eg. AlPO4, Al(OH)3, QIL A,
     QS21, Ca3(PO4)2, Ca(OH)2, Zn(OH)2 a glycolipid analogue or an octadecyl ester of an amino acid. The OMPs of Haemophilus strains may be purified
     using eg. DEAE-Sephacel (RTM) and hydroxyapatite columns.
           USE - The ICMs can be used in vaccines to confer protection against
     disease caused by the Streptococcus strain and the Haemophilus strain
     (claimed). The ICMs and antibodies produced using the ICMs can also be
     used in diagnostic procedures and kits.
          ADVANTAGE - The ICMs provide an enhanced immune response to CPs of
     Streptococcus strains without the necessity to employ carrier proteins.
     They also provide an immune response to the OMP of the Haemophilus strain.
     Dwg.0/8
FS
     CPI EPI
     CPI: B04-C02F; B04-G07; B04-N03; B11-C07A; B12-K04A4; B14-A01A;
          B14-A01B2; B14-S11B; D05-H04; D05-H07; D05-H10
     EPI: S03-E14H4
    ANSWER 27 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
L54
     1994-200269 [24]
                         WPIX
AN
DNC
    C1994-091569
     Nucleic acid encoding D15 outer membrane protein - especially of Haemophilus
     influenzae, and related proteins, vectors, antisera etc. useful in
```

vaccines, for diagnosis and for passive immunisation...

```
DC
     B04 D16
IN
     CHONG, P; KLEIN, M; LOOSMORE, S; SIA, D Y C; THOMAS,
     W; YANG, Y; LOOSMORE, S M; SIA, D; YANG, Y P
     (CONN-N) CONNAUGHT LAB LTD; (AVET) AVENTIS PASTEUR LTD
PA
CYC
     28
PΤ
     WO 9412641
                     A1 19940609 (199424)*
                                                       C12N015-31
        RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL PT SE
         W: AU BR CA FI JP KR NO NZ RU UA US
     AU 9455565
                     A 19940622 (199436)
                                                       C12N015-31
     EP 668916
                     A1 19950830 (199539)
                                                       C12N015-31
         R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE
                     W 19960319 (199644)
B 19971113 (199803)
     JP 08502417
                                                156
                                                       C12N015-09
     AU 683435
                                                       C12N015-31
     BR 9307510
                     A 19990601 (199927)
                                                       C12N015-31
     JP 2907552
                     B2 19990621 (199930)
                                                181
                                                       C12N015-09
                     A 20000111 (200010)
     US 6013514
                                                       A61K039-102
     US 6083743
                     A 20000704 (200036)
                                                       A61K039-102
                                                                       <---
                     C1 19991120 (200041)
                                                       C12N015-31
     RU 2141528
     KR 216390
                     B1 19990816 (200104)
                                                       C12N015-31
     US 6264954
                     B1 20010724 (200146)
                                                       A61K039-102
ADT WO 9412641 A1 WO 1993-CA501 19931123; AU 9455565 A AU 1994-55565 19931123;
     EP 668916 A1 WO 1993-CA501 19931123, EP 1994-900671 19931123; JP 08502417
     W WO 1993-CA501 19931123, JP 1994-512608 19931123; AU 683435 B AU
     1994-55565 19931123; BR 9307510 A BR 1993-7510 19931123, WO 1993-CA501
     19931123; JP 2907552 B2 WO 1993-CA501 19931123, JP 1994-512608 19931123;
     US 6013514 A WO 1993-CA501 19931123, US 1995-433522 19950912; US 6083743 A
     Cont of WO 1993-CA501 19931123, Cont of US 1995-433522 19950912, US
     1998-135166 19980818; RU 2141528 C1 WO 1993-CA501 19931123, RU 1995-117238
     19931123; KR 216390 B1 WO 1993-CA501 19931123, KR 1995-702081 19950523; US
     6264954 B1 Cont of WO 1993-CA501 19931123, Div ex US 1995-433522 19950912,
     US 1997-942046 19971001
FDT AU 9455565 A Based on WO 9412641; EP 668916 Al Based on WO 9412641; JP
     08502417 W Based on WO 9412641; AU 683435 B Previous Publ. AU 9455565,
     Based on WO 9412641; BR 9307510 A Based on WO 9412641; JP 2907552 B2
     Previous Publ. JP 08502417, Based on WO 9412641; US 6013514 A Based on WO
     9412641; RU 2141528 Cl Based on WO 9412641
PRAI GB 1992-24584
                          19921123
REP 01Jnl.Ref; EP 281673; EP 378929; US 5013664; WO 9106652
     ICM A61K039-102; C12N015-09; C12N015-31
IC
         A61K039-12; A61K039-395; C07H021-04; C07K013-00;
          C07K014-11; C07K014-195; C07K014-285; C07K016-12; C12N015-62
ICA C12P021-02; G01N033-569
ICI C12N015-31, C12R001:21; C12P021-02, C12R001:19; C12N015-31, C12R001:21
         9412641 A UPAB: 19940803
     New nucleic acid (I) contains at least a portion coding for a D15 outer
     membrane protein (omp) and has a sequence which is (a) any of 5 (all about
     3000bp) reproduced in the specification, or complementary sequences or (b)
     hybridsable under stringent conditions with such sequences. Also new are
     (1) recombinant plasmids containing a segment of (I) at least 18 bp long (and
     opt. expression control elements, (12) proteins (II) encoded by these plasmids; (3) purified D15 omp (III); (4) synthetic polypeptides with
     sequences corresp. to (II) or (III), or their variants and mutants which
     retain immunogenicity; (5) antisera or antibodies specific for (II), (III)
     or immunologous containing them; (6) chimeric molecules consisting of (II) or
     (III) bonded to another polypeptides, protein or polysaccharides.
          USE - (I), (II) and the synthetic polypeptides are useful in vaccines
     to protect against Haemophilus. D15 can also be used as a carrier for
     polysaccharide antigens to form conjugate vaccines against other bacteria;
     to induce immunity to abnormal polysaccharides or tumour cells and to
     generate anti-tumour antibodies, for coupling to toxins etc. (I), (II)
     synthetic peptides and antisera can also be used diagnostically (in
     hybridisation or immunoassay procedures) and antibodies can be used for
    passive immunisation.
     Dwg.0/11
FS
     CPI
FA
    AB
     CPI: B04-C01; B04-C02; B04-E02F; B04-E08; B04-G01; B04-N0300E;
          B04-N04; B12-K04; B14-A01A; B14-H01B; B14-S11A;
          B14-S11B; D05-H11; D05-H12A; D05-H12E; D05-H17A; D05-H17C
    ANSWER 28 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
AN
    1993-134388 [16]
                        WPIX
DNC
    C1993-059997
ΤI
     Oligoside derived from antigenic polyoside of a pathogen - useful for
     treating and preventing e.g. bacterial infections and mycoses.
DC
```

```
MOREAU, M
IN
     (AVET) AVENTIS PASTEUR; (INMR) PASTEUR MERIEUX SERUMS & VACCINS
PA
     SA; (INMR) PASTEUR MERIEUX SERUMS & VACCINS
CYC
     WO 9307178
                      A1 19930415 (199316) * EN 38
                                                           C08B037-00
ΡI
        RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL SE
         W: AU CA FI HU JP KR NO US
                      A1 19930416 (199328)
                                                    29
                                                           C08B037-00
     FR 2682388
     AU 9229469
                      A 19930503 (199334)
     FI 9302626
                      A 19930609 (199334)
                                                           C08B000-00
                      A1 19930929 (199339)
     EP 562107
                                              FR
         R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL SE
     NO 9302102
                      A 19930805 (199343)
                      W 19940714 (199432)
                                                           A61K039-02
     JP 06506233
                      B 19950713 (199535)
T 19950928 (199546)
     AU 661071
                                                           C08B037-00
     HU 70298
     US 6007818
                      A 19991228 (200007)
                                                           A61K039-00
                                                                           <-- .
     NO 306905
                      B1 20000110 (200009)
                                                           C08B037-00
     US 6045805
                      A 20000404 (200024)
                                                           A61K039-085
     KR 249709
                      B1 20000315 (200122)
                                                           C08B037-00
                                                           C08B037-00
     HU 219672
                      B 20010628 (200143)
                      B1 20020502 (200230)
                                             FR
                                                           C08B037-00
     EP 562107
         R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL SE
     DE 69232585
                      E 20020606 (200245)
                                                           C08B037-00
                                                           C08B037-00
                      T3 20021116 (200302)
     ES 2174839
     FI 110164
                      B1 20021213 (200306)
                                                           A61K039-02
                      C 20030429 (200337)
     CA 2098105
                                             FR
                                                           C08B037-00
    WO 9307178 A1 WO 1992-FR955 19921009; FR 2682388 A1 FR 1991-12478 19911010; AU 9229469 A AU 1992-29469 19921009; FI 9302626 A WO 1992-FR955
     19921009, FI 1993-2626 19930609; EP 562107 A1 EP 1992-923831 19921009, WO
     1992-FR955 19921009; NO 9302102 A WO 1992-FR955 19921009, NO 1993-2102
     19930609; JP 06506233 W WO 1992-FR955 19921009, JP 1993-506690 19921009;
     AU 661071 B AU 1992-29469 19921009; HU 70298 T WO 1992-FR955 19921009, HU
     1993-1682 19921009; US 6007818 A Div ex WO 1992-FR955 19921009, Div ex US
     1993-70446 19931007, US 1995-474190 19950607; NO 306905 B1 WO 1992-FR955
     19921009, NO 1993-2102 19930609; US 6045805 A Cont of WO 1992-FR955 19921009, Cont of US 1993-70446 19931007, US 1995-474194 19950607; KR
     249709 B1 WO 1992-FR955 19921009, KR 1993-701727 19930609; HU 219672 B WO
     1992-FR955 19921009, HU 1993-1682 19921009; EP 562107 B1 EP 1992-923831
     19921009, WO 1992-FR955 19921009; DE 69232585 E DE 1992-632585 19921009,
     EP 1992-923831 19921009, WO 1992-FR955 19921009; ES 2174839 T3 EP 1992-923831 19921009; FI 110164 B1 WO 1992-FR955 19921009, FI 1993-2626
     19930609; CA 2098105 C CA 1992-2098105 19921009, WO 1992-FR955 19921009
FDT AU 9229469 A Based on WO 9307178; EP 562107 Al Based on WO 9307178; JP
     06506233 W Based on WO 9307178; AU 661071 B Previous Publ. AU 9229469,
     Based on WO 9307178; HU 70298 T Based on WO 9307178; NO 306905 B1 Previous
     Publ. NO 9302102; HU 219672 B Previous Publ. HU 70298, Based on WO
     9307178; EP 562107 B1 Based on WO 9307178; DE 69232585 E Based on EP
     562107, Based on WO 9307178; ES 2174839 T3 Based on EP 562107; FI 110164
     Bl Previous Publ. FI 9302626; CA 2098105 C Based on WO 9307178
PRAI FR 1991-12478
                            19911010
    3.Jnl.Ref; BE 1000118; EP 245045; EP 97407
     ICM A61K039-00; A61K039-02; A61K039-085;
         C08B000-00; C08B037-00
A61K031-70; A61K031-715; A61K035-74; A61K038-00; A61K039-08
           ; A61K039-09; A61K039-102; A61K039-106;
          A61K039-112; C07H003-06; C07K001-00; C07K014-00
AB
          9307178 A UPAB: 19930924
     New oligoside (I), retaining at least one antigenic determinant of an
     antigenic polyoside (A) derived from a pathogen, is prepared by (1)
     oxido-reductive depolymerisation of (A); (2) recovering (I) and opt. (3)
     coupling it to a conjugation partner or to a carrier to form a conjugate.
     Pref. (I) have mean elution constant on Sepharose 4BCL of 0.2-1, best
     0.6-0.7 (equivalent to mol.weight 30000-60000, dextran equivalent) and is derived
     from the capsular polysaccharides of a pathogenic Staphylococcus,
     Streptococcus, Klebsiella, Salmonella, Escherichia, Neisseria or
     Haemophilus, especially Salmonella typhi, Strep. pneumoniae, N. meningitidis or
     H. influenzae.
     (I) is pref. conjugated to a peptide, protein or organic polymer, most pref. pertussis, cholera, tetanus or diphtheria toxin.4
     Alternatively, (I) is incorporated into a vector which stimulates
     immunogenicity in mammals, or into a liposome.

USE/ADVANTAGE - (I), especially in conjugated form, are useful in vaccines
     to protect against (or reduce the effects of) bacterial infections or
     mycoses. The usual dose (by any standard route) is 1-200 microg in 0.5ml.
```

This method of (I) production produces fragments of homogeneous size which

Graser 10/030313

Page 101

```
retain the essential structural determinants; is simple and inexpensive,
     and can be applied to any polyoside structure.
     0/6
FS
     CPI
     AB
FA
     CPI: B02-V02; B04-B02B1; B04-C02F; B12-A01; B12-A02C
     ANSWER 29 OF 29 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
L54
AN
     1991-178107 [24]
                          WPIX
DNC
     C1991-076917
     New mammalian cytokine interleukin-11 - for use in treating immune system
TI
     disorders, e.g. deficiencies in haematopoietic progenitor or stem cells,
     and cancer.
DC
     BENNETT, F K; PAUL, S R; YANG, Y; STEPHEN, R P; YANG, Y C (CHIL-N) CHILDRENS MEDICAL CENT; (GEMY) GENETICS INST INC; (CHIL-N)
IN
PA
     CHILDRENS MED CENTER CORO
CYC
     21
     WO 9107495
                       A 19910530 (199124) *
PΙ
         RW: AT BE CH DE DK ES FR GB GR IT LU NL SE
         W: AU CA HU JP KR US
     AU 9067578
                       A 19910613 (199137)
     EP 504177
                       A1 19920923 (199239)
                                               EN
                                                     69
                                                           C12N015-24
         R: AT BE CH DE DK ES FR GB GR IT LI LU NL SE
                       A 19920722 (199242)
                                                            C07K013-00
     DK 9200666
     US 5215895
                       A 19930601 (199323)
                                                     17
                                                            C12P021-02
                       W 19930715 (199333)
     JP 05504560
                                                     69
                                                            C07K013-00
     AU 644389
                       B 19931209 (199405)
                                                            C07K013-00
                       T 19940128 (199409)
     HU 64595
                                                           C12N015-24
     US 5371193
                       A 19941206 (199503)
                                                     18
                                                            C07K013-00
     JP 2688539
                       B2 19971210 (199803)
                                                     20
                                                            C07K014-54
                      A 19971223 (199806)
     US 5700664
                                                     20
                                                           C12N015-24
                      A 19980113 (199812)
B 19970430 (199821)
     JP 10004982
                                                     21
                                                           C12N015-09
     MX 184567
                                                            C12N015-024
     JP 2783361
                       B2 19980806 (199836)
                                                     21
                                                            C12N015-09
     HU 215233
                       B 19981130 (199903)
                                                            C12N015-24
                       A 19981229 (199908)
     US 5854028
                                                           C12N015-63
     KR 9705050
                      B1 19970411 (199938)
                                                           C12N015-24
     US 6066317
                       A 20000523 (200032)
                                                           A61K038-19
                       B1 20010207 (200109) EN
     EP 504177
                                                           C12N015-24
         R: AT BE CH DE DK ES FR GB GR IT LI LU NL SE
     DE 69033700
                      E 20010315 (200122)
                                                           C12N015-24
     CA 2069428
                       C 20010731 (200147)
                                               EN
                                                            C12N015-24
     ES 2156852
                       T3 20010801 (200149)
                                                           C12N015-24
ADT EP 504177 A1 EP 1990-917548 19901120, WO 1990-US6803 19901120; DK 9200666
     A WO 1990-US6803 19901120, DK 1992-666 19920521; US 5215895 A CIP of US
     1989-441100 19891122, US 1990-526474 19900521; JP 05504560 W WO
     1990-US6803 19901120, JP 1991-500597 19901120; AU 644389 B AU 1990-67578
     19901120; HU 64595 T WO 1990-US6803 19901120, HU 1992-1712 19901120; US 5371193 A Div ex US 1990-526474 19900521, US 1993-17522 19930212; JP
     2688539 B2 WO 1990-US6803 19901120, JP 1991-500597 19901120; US 5700664 A
     CIP of US 1989-441100 19891122, CIP of US 1990-526474 19900521, WO
     1990-US6803 19901120, US 1992-949516 19921119; JP 10004982 A Div ex JP
     1991-500597 19901120, JP 1997-51468 19901120; MX 184567 B MX 1992-3439 19920626; JP 2783361 B2 Div ex JP 1991-500597 19901120, JP 1997-51468
     19901120; HU 215233 B WO 1990-US6803 19901120, HU 1992-1712 19901120; US
     5854028 A CIP of US 1989-441100 19891122, CIP of US 1990-526474 19900521,
     Div ex US 1992-949516 19921119, US 1997-814459 19970310; KR 9705050 B1 WO
     1990-US6803 19901120, KR 1992-701213 19920522; US 6066317 A CIP of US 1989-441100 19891122, Cont of US 1990-526474 19900521, Div ex US
     1992-949516 19921119, Cont of US 1997-814459 19970310, US 1998-122525
     19980724; EP 504177 B1 EP 1990-917548 19901120, WO 1990-US6803 19901120;
     DE 69033700 E DE 1990-633700 19901120, EP 1990-917548 19901120, WO
     1990-US6803 19901120; CA 2069428 C CA 1990-2069428 19901120, WO
     1990-US6803 19901120; ES 2156852 T3 EP 1990-917548 19901120
FDT EP 504177 A1 Based on WO 9107495; JP 05504560 W Based on WO 9107495; AU
     644389 B Previous Publ. AU 9067578, Based on WO 9107495; HU 64595 T Based
     on WO 9107495; US 5371193 A Div ex US 5215895; JP 2688539 B2 Previous
     Publ. JP 05504560, Based on WO 9107495; US 5700664 A CIP of US 5215895
     Based on WO 9107495; JP 2783361 B2 Previous Publ. JP 10004982; HU 215233 B
     Previous Publ. HU 64595, Based on WO 9107495; US 5854028 A CIP of US
     5215895, Div ex US 5700664; US 6066317 A Cont of US 5215895, Div ex US 5700664, Cont of US 5854028; EP 504177 B1 Based on WO 9107495; DE 69033700
     E Based on EP 504177, Based on WO 9107495; CA 2069428 C Based on WO
     9107495; ES 2156852 T3 Based on EP 504177
```

19901120; US 1989-441100

PRAI WO 1990-US6803

19891122:

Graser 10/030313 Page 102

```
WO 1990-US6803U
                         19901120; US 1993-17522
                                                         19930212;
                          19921119; US 1997-814459
     US 1992-949516
                                                         19970310;
    US 1998-122525
                          19980724
    2.Jnl.Ref
REP
    ICM A61K038-19; C07K013-00; C12N015-024; C12N015-09; C12N015-24;
         C12N015-63; C12P021-02
         A61K037-02; A61K037-66; A61K038-20; A61K039-395;
         C07H015-12; C07K013-000; C07K014-00; C07K015-06; C12N001-21;
         C12N015-19; C12P021-002
ICA
    A61K038-00; C07H021-04; C07K014-54; C12N005-10
    A61K038:00; A61K037-02, A61K037:66; C12P021-02, C12R001:91; C12P021-02,
ICI
         C12R001:91
AB
         9107495 A UPAB: 19940329
    Mammalian IL-11 (I), free from other proteins, is new. (I) has a molecular
    weight of 20 kD (SDS-PAGE and calculations) and biological activity in T1165
    assays, megakaryoctye colony forming assays with IL-3 and B cell plaque
     forming assays. Also new are a process for producing (I) recombinantly,
     DNA encoding (I), a cell transformed with this DNA, a plasmid vector
    containing the DNA and homogeneous mammalian (I) having biological activity in
     the T1165 assay without IL-6. Also present in the compsn. may be other
     cytokine e.g. IL-1 to IL-9, GM-CSF, G-CSF, M-CSF, the interferons,
     Meg-CSF, MIF, LIF, TNF and erythropoietin, haematopoietins e.g. IL-3 or
     IL-6, growth factors or antibodies. Dosage of (I) is 1-1000 micro-g or
     50-5000 units, where a unit is the concentration leading to half maximal
     stimulation in the T1165 assay.
         USE/ADVANTAGE - (I) is used to stimulate or treat disorders of the
     immune system e.g. deficiencies in haematopoietic progenitor stem cells
     (e.g. following bone marrow transplantation), and to treat cancer and
     other pathological states caused by disease, exposure to radiation or
     drugs and e.g. leukopenia, bacterial and viral infections, anaemia and B
     or T cells deficiencies. (I) is also used to prolong the effects of
     vaccines. Use of (I) does not create undesirable side effects. @(69pp
    Dwg.No.0/2)
     0/2
FS
    CPI
    AB
FA
    CPI: B04-B02B1; B04-B04A1; B04-C01G; B12-A01; B12-A06; B12-A07;
MC
         B12-G05; B12-G07; B12-H01; D05-C12; D05-H12
=> d all 156 tot
    ANSWER 1 OF 4 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
1.56
     2000-594517 [56] WPIX
AN
     2000-594515 [56]; 2000-594516 [56]; 2000-679550 [66]; 2001-006956 [01]
CR
    C2000-177617
DNC
    A Streptococcus pneumoniae vaccine for preventing pneumonia and meningitis
     comprises a polysaccharide antigen conjugated to protein D from
    Haemophilus influenzae.
DC
    B04 D16
    CAPIAU, C; DESCHAMPS, M; DESMONS, P M; LAFERRIERE, C A J; POOLMAN, J;
IN
    PRIEELS, J; POOLMAN, J P J
     (SMIK) SMITHKLINE BEECHAM BIOLOGICALS
PA
   93
CYC
PΙ
    WO 2000056360
                   A2 20000928 (200056) * EN 77
                                                      A61K039-385
       RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL
            OA PT SD SE SL SZ TZ UG ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ
            EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK
            LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI
            SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
    AU 2000034307 A 20001009 (200103)
                                                      A61K039-385
    BR 2000009163
                    A 20011226 (200206)
                                                      A61K039-385
                                                                     <--
    EP 1163000
                    A2 20011219 (200206) EN
                                                      A61K039-385
        R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
            RO SE SI
    CZ 2001003380
                    A3 20020313 (200223)
                                                      A61K039-385
                                                                     <--
    KR 2002000549
                    A 20020105 (200244)
                                                      A61K039-385
                                                                     <--
                                                                     <--
                       20020528 (200249)
    HU 2002000367
                    В
                                                      A61K039-385
    CN 1351503
                    A 20020529 (200258)
                                                      A61K039-385
                                                                     <--
                    B 20020801 (200261)
                                                      A61K039-385
    AU 750913
                                                                     <--
    ZA 2001007637
                    A 20020828 (200264)
                                                97
                                                      A61K000-00
    JP 2002540075
                    W 20021126 (200307)
                                                96
                                                      A61K039-09
    MX 2001009455
                    A1 20020301 (200362)
                                                      A61K039-005
                                                                     <--
                    A 20040227 (200418)
    NZ 513840
                                                      A61K039-385
                                                                     <--
```

```
ADT WO 2000056360 A2 WO 2000-EP2468 20000317; AU 2000034307 A AU 2000-34307
     20000317; BR 2000009163 A BR 2000-9163 20000317, WO 2000-EP2468 20000317;
    EP 1163000 A2 EP 2000-912626 20000317, WO 2000-EP2468 20000317; CZ 2001003380 A3 WO 2000-EP2468 20000317, CZ 2001-3380 20000317; KR 2002000549 A WO 2000-EP2468 20000317, KR 2001-711939 20010919; HU
     2002000367 B WO 2000-EP2468 20000317, HU 2002-367 20000317; CN 1351503 A
     CN 2000-807528 20000317; AU 750913 B AU 2000-34307 20000317; ZA 2001007637
     A ZA 2001-7637 20010917; JP 2002540075 W JP 2000-606264 20000317, WO
     2000-EP2468 20000317; MX 2001009455 A1 WO 2000-EP2468 20000317, MX
     2001-9455 20010919; NZ 513840 A NZ 2000-513840 20000317, WO 2000-EP2468
     20000317
FDT AU 2000034307 A Based on WO 2000056360; BR 2000009163 A Based on WO
     2000056360; EP 1163000 A2 Based on WO 2000056360; CZ 2001003380 A3 Based
     on WO 2000056360; KR 2002000549 A Based on WO 2000056360; HU 2002000367 B
     Based on WO 2000056360; AU 750913 B Previous Publ. AU 2000034307, Based on
     WO 2000056360; JP 2002540075 W Based on WO 2000056360; MX 2001009455 A1
     Based on WO 2000056360; NZ 513840 A Based on WO 2000056360
PRAI GB 1999-16677
                          19990715; GB 1999-6437
                                                           19990319:
                          19990420; GB 1999-9466
     GB 1999-9077
TC
    ICM A61K000-00; A61K039-005; A61K039-09;
          A61K039-385
     ICS A61K035-74; A61K039-02; A61K039-04;
          A61K039-085; A61K039-095; A61K039-102;
          A61K039-112; A61K039-116; A61K039-39;
          A61P011-00; A61P031-04; C07K014-285
    C12N015-09
ICA
     WO 200056360 A UPAB: 20040316
AB
     NOVELTY - A polysaccharide conjugate antigen (I) comprising a
     polysaccharide antigen derived from a pathogenic bacterium conjugated to
     protein D (or a fragment) from Haemophilus influenzae, is new.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the
     following:

    an immunogenic composition comprising (I);

          (2) an immunogenic composition comprising Neisseria meningitidis
     protein D polysaccharide conjugate antigen;
          (3) an immunogenic composition comprising Haemophilus
     influenzae b protein D polysaccharide conjugate antigen;
          (4) an immunogenic composition comprising conjugated capsular
     polysaccharides of Streptococcus pneumoniae, Haemophilus
     influenzae b , meningococcus C and meningococcus Y, the carrier protein
     for at least one of the polysaccharides is protein D from H. influenzae;
          (5) a vaccine comprising (1)-(4); and
          (6) a method for producing an immunogenic composition to a pathogenic
     bacterium comprising:
          (a) isolating a polysaccharide antigen from a pathogenic bacterium;
          (b) activating the polysaccharide; and
          (c) conjugating the polysaccharide to protein D.
          ACTIVITY - Antibacterial. No biological data given
          MECHANISM OF ACTION - Vaccine.
          USE - The bacterial polysaccharide antigen vaccines are used to
     induce an immune response to Streptococcus pneumoniae and is used to
     prevent pneumonia, bacteremia, meningitis and acute otitis media.
          ADVANTAGE - The conjugation of the antigen to a larger immunogenic
     protein increases the induced immune response, especially in children less
     than two years old.
     Dwg.0/3
FS
     CPI
     AB; DCN
FA
     CPI: B04-B04C1; B04-C02F; B04-C02V; B04-F10A; B04-F10B; B04-N03;
MC
          B04-N05; B04-N06; B12-M07; B14-A01B2; B14-S11B; D05-H07
L56
    ANSWER 2 OF 4 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
     1994-025891 [03]
                        WPIX
AN
     1992-249778 [30]
CR
DNC
    C1994-011927
     New adhesion-oligosaccharide conjugate - useful as vaccine for Haemophilus
     influenzae, and new synthetic poly ribosyl ribotol phosphate
     oligosaccharide(s).
DC
     B04 D16
     KRIVAN, H C; NORBERG, N T; SAMUELS, J E; SAMUEL, J E
IN
     (MICR-N) MICROCARB INC; (ANTE-N) ANTEX BIOLOGICS FORMERLY MICROCARB INC;
PA
     (ANTE-N) ANTEX BIOLOGICS INC; (ANTE-N) ANTEXBIOLOGICS INC
CYC
                     A1 19940106 (199403) * EN 124
                                                        A61K039-00
ΡI
        RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL PT SE
         W: CA JP US
```

```
A1 19950412 (199519) EN
     EP 647139
                                                           A61K039-00
         R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE
                   W 19951026 (199551)
A 19971021 (199748)
A4 19970903 (199815)
     JP 07509693
                                                    33
                                                           C07K014-11
     US 5679547
                                                    35
                                                           C12P021-00
     EP 647139
                                                           A61K039-00
     US 5721115
                      A 19980224 (199815)
                                                    36
                                                           C12P021-06
     JP 2805174
                      B2 19980930 (199844)
                                                    42
                                                           C07K014-11
                      A 19981201 (199904)
     US 5843463
                                                           A61K039-102
                      C 19990907 (200003) EN
                                                           C12N015-31
     CA 2138765
ADT WO 9400149 A1 WO 1993-US6016 19930622; EP 647139 A1 EP 1993-916717
     19930622, WO 1993-US6016 19930622; JP 07509693 W WO 1993-US6016 19930622,
     JP 1994-502559 19930622; US 5679547 A CIP of WO 1986-DK14 19860206, CIP of US 1990-631698 19901221, CIP of US 1991-810966 19911220, Div ex US
     1992-903079 19920622, US 1995-485569 19950607; EP 647139 A4 EP 1993-916717
     19930622; US 5721115 A CIP of US 1990-631698 19901221, CIP of US
     1991-810966 19911220, Div ex US 1992-903079 19920622, US 1995-480993
     19950607; JP 2805174 B2 WO 1993-US6016 19930622, JP 1994-502559 19930622; US 5843463 A CIP of US 1990-631698 19901221, CIP of US 1991-810966
     19911220, US 1992-903079 19920622; CA 2138765 C CA 1993-2138765 19930622,
     WO 1993-US6016 19930622
FDT EP 647139 Al Based on WO 9400149; JP 07509693 W Based on WO 9400149: JP
     2805174 B2 Previous Publ. JP 07509693, Based on WO 9400149; CA 2138765 C
     Based on WO 9400149
PRAI US 1992-903079
                            19920622; WO 1986-DK14
                                                              19860206:
                            19901221; US 1991-810966
19950607; US 1995-480993
     US 1990-631698
                                                              19911220:
     US 1995-485569
                                                              19950607
     2.Jnl.Ref; EP 276516; EP 320942; US 4455296; 3.Jnl.Ref; EP 338265; WO
REP
     9210936
     ICM A61K039-00; A61K039-102; C07K014-11; C12N015-31;
IC
           C12P021-00; C12P021-06
          A61K037-02; A61K038-00; A61K039-02; C07H001-04; C07H013-00;
           C07H015-02; C07H021-02; C07H021-04; C07K003-28; C07K014-00;
          C07K014-285; C07K015-00; C07K019-00; C12N001-21; C12N015-06; C12N015-09; C12P017-00; C12P021-02; C12P021-04; C12Q001-34;
          G01N033-53
           9400149 A UPAB: 20000118
AB
     An immunogenic oligosaccharide-protein conjugate (I) comprising a
     polyribosyl-ribotol phosphate (PRP) fragment coupled to an Haemophilus
     influenzae adhesin protein (II) is new.
           (II) is pref. an H. influenzae outer membrane protein with a mol.weight
     of about 47000 daltons, and purified (II) is claimed per se.

USE - (I), as well as their protein components, may be used in
     vaccines against both invasive and non-invasive strains of H. influenzae.
     (I), (II) and oligomers are also useful as reagents for scientific
     research on the properties of pathogenicity, virulence and infectivity of
     H. influenzae, as well as host defence mechanisms. E.g. the novel DNA can
     be used in an oligonucleotide probe to identify the DNA of other
     microorganisms which might encode an adhesion for such organism. (I) can
     be used to prepare a monoclonal antibody useful to further purity compsns.
     containing (II) by affinity chromatography. (II) could also be applied to
     standard immunoassays to screen for the presence of antibodies to H.
     influenza in a sample. (IV) are intermediates in the synthesis of (III),
     which may be used to prepare (Ia).
     Dwg.0/9
     CPI
FS
     AB; DCN
FΑ
MC
     CPI: B04-C02; B04-C02X; B04-E02F; B04-E03F; B04-E08; B04-N03; B05-B01M;
          B12-K04; B14-A01A; B14-S11B; D05-H07; D05-H12;
          D05-H12E
L56 ANSWER 3 OF 4 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
     1990-132245 [17] WPIX
DNC
     C1990-058101
ΤI
     Capsular polysaccharide adhesion antiqen - from coaqulase negative
     bacteria used to prevent or treat infection caused by staphylococcal
     strains.
DC
     A96 B04 D16 D22
IN
     PIER. G B
     (BGHM) BRIGHAM & WOMENS HOSPITAL; (BGHM) BRIGHAM & WOMENS HOSPITAL INC;
PA
     (BRIG-N) BRIGHAM & WOMENS HO; (PIER-I) PIER G B
CYC
     15
PΙ
     WO 9003398
                      A 19900405 (199017)*
        RW: AT BE CH DE FR GB IT LU NL SE
         W: AU JP
                      A 19900418 (199027)
     AU 8943430
     EP 436648
                      A 19910717 (199129)
```

```
R: AT BE CH DE FR GB IT LI LU NL SE
                     A 19911008 (199143)
W 19920326 (199219)
     US 5055455
     JP 04501718
                                                   15
                      C 19930504 (199323)
A4 19911113 (199520)
                                                          C12P019-04
     CA 1317288
     EP 436648
     US 5980910
                      A 19991109 (199954)
                                                          A61K039-09
     US 6399066
                      B1 20020604 (200242)
                                                         A61K039-40
                                                                          <--
                    A1 20020926 (200265)
     US 2002136730
                                                          A61K039-40
                                                                          e - -
     US 6743431
                     B2 20040601 (200436)
                                                         A61K039-085
                                                                          <--
ADT EP 436648 A EP 1989-911517 19890928; US 5055455 A US 1988-250417 19880928;
     JP 04501718 W JP 1989-510684 19890928; CA 1317288 C CA 1989-614255
     19890928; EP 436648 A4 EP 1989-911517
                                                      ; US 5980910 A Div ex US
     1988-250417 19880928, Cont of US 1991-727982 19910710, Cont of US
     1993-33756 19930318, US 1994-336688 19941107; US 6399066 B1 Div ex US
     1988-250417 19880928, Cont of US 1991-727982 19910710, Cont of US 1993-33756 19930318, Div ex US 1994-336688 19941107, US 1999-393832
     19990910; US 2002136730 Al Div ex US 1988-250417 19880928, Cont of US
     1991-727982 19910710, Cont of US 1993-33756 19930318, Div ex US
     1994-336688 19941107, Div ex US 1999-393832 19990910, US 2002-93582
     20020308; US 6743431 B2 Div ex US 1988-250417 19880928, Cont of US
     1991-727982 19910710, Cont of US 1993-33756 19930318, Div ex US
     1994-336688 19941107, Div ex US 1999-393832 19990910, US 2002-93582
     20020308
FDT US 5980910 A Div ex US 5055455; US 6399066 B1 Div ex US 5055455, Div ex US
     5980910; US 2002136730 A1 Div ex US 5055455, Div ex US 5980910, Div ex US
     6399066; US 6743431 B2 Div ex US 5055455, Div ex US 5980910, Div ex US
     6399066
PRAI US 1988-250417
                           19880928; US 1991-727982
                                                             19910710:
                           19930318; US 1994-336688
     US 1993-33756
                                                             19941107:
     US 1999-393832
                           19990910; US 2002-93582
                                                             20020308
REP
     US 4789735; US 4830852; 2.Jnl.Ref; EP 302781; FR 2410043
     A61K037-00; A61K039-02; A61K039-08; C07K015-04;
     C07K015-14; C12P021-00
     ICM A61K039-085; A61K039-09; A61K039-40;
          C12P019-04
     ICS A61K037-00; A61K039-02; A61K039-08; C07K015-04; C07K015-14; C08B037-00; C12P021-00
AB
          9003398 A UPAB: 19991122
     The following are m (A) a capsular polysaccharide adhesion from
     coaqulase-negative bacteria (e.q. Staphylococcus epidermidis or hominus
     strains) in pure form; (B) a vaccine against coagulase-negative
     staphylococci comprising a vehicle containing the pure capsular polysaccharide
     adhesionm antigen specific to the staphylocci; the vehicle may be e.g.
     Freund's complete or incomplete adjuvant, saline, serum albumin or
     saponin; (C) monovalent antibody (MAb) against capsular polysaccharide
     adhesin of coagulase-negative bacteria.
          USE/ADVANTAGE - The polysaccharide adhesin can produce
     antibodies which prevent the adherence of adhesin-bearing
     pathogenic bacteria to the recipients tissue cells or polymeric medical
     prostheses or catheters and can therefore be used for preventing or
     treating diseases and infections due to staphylococci. The adhesin
     can also be used to screen polymeric materials for resistance to attachment by bacteria. The MAbs can be administered to prevent or reduce
     infections by coagulase-negative staphylococci. The adhesin
     -specific antibodies can also be used in affinity chromatography and in
     diagnosis and assays.
     Dwg.0/5
FS
     CPI
     AΒ
FΔ
MC
     CPI: A09-C; A12-V01; A12-V02; B02-V02; B04-B04C5;
          B04-C02; B12-A01; D05-H07; D05-H11
L56 ANSWER 4 OF 4 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
AN
     1983-42112K [18]
                         WPIX
                         DNC C1983-041031
DNN
    N1983-076352
     Mono clonal antibodies against bacterial adhesion(s) - useful for treating
     diarrhoea in neonates, respiratory diseases and burns.
DC
     B04 D16 S03
IN
     SADOWKI, P L
     (MOLE-N) MOLECULAR GENETICS INC
PA
CYC 12
                      A 19830427 (198318)* EN
PΙ
     EP 77734
                                                   37
         R: BE DE FR GB IT LU NL
     AU 8289454
                      A 19830428 (198324)
                      A 19830613 (198329)
A 19830620 (198331)
     JP 58099423
     DK 8204621
```

```
US 4443549
                     A 19840417 (198418)
                     A 19850528 (198526)
A 19870324 (198714)
     CA 1187822
     US 4652448
     US 4443549 A US 1982-428622 19821007; US 4652448 A US 1983-558518 19831206
ADT
PRAI US 1981-312993
                           19811019; US 1982-428622
                                                           19821007:
     US 1983-558518
                           19831206
     4.Jnl.Ref; No-SR.Pub
REP
     A61K039-40; C12N005-00; C12N015-00; C12P001-00; C12R001-91;
IC
     G01N033-54
     EΡ
            77734 A UPAB: 19930925
     Production of anti-adhesion antibodies comprises fusion of a cell producing
     the antibodies with a myeloma to provide a fused cell hybrid, followed by
     propagation of the hybrid and collection of the antibodies.
          Production of antipilus antibodies comprises injection of a BALB/C mouse
     with a bacterial pilus to induce formation of antibacterial pilus
     antibody-producing cells of the mouse. Then a fused cell hybrid of the
     cells is produced with P3/NSI/1-Ag4-1 myeloma cells, and the hybrid is
     cultured in vitro in selective HAT medium, isolated and propagated and the
     resulting antibodies are harvested.
          Continuous cell line producing anti-adhesion antibodies and
     comprising a fused cell hydrid of an anti-adhesion antibody-producing cell
     and a myeloma cell is new, and cell line 2BD4E4 (ATCC HB8178) is new.
          Monoclonal antibodies against Escherichia coli adhesions are useful
     for admin. to animals and humans, especially for the prophylaxis and treatment
     of enterotoxigenic diarrhoeal diseases in neonatal calves, lambs and
     piglets. The antibodies are obtainable in lage amts. and are useful as
     highly sensitive and specific probes in medicinal and veterinary
     diagnosis, etc. The anti-adhesion antibodies may also be useful against
     respiratory diseases and burn infections and against other bacterial
     diseases. The antibodies are also useful in affinity chromatography
     systems and in the assay of adhesions.
FS
     CPI EPI
FA
     AB
MC
     CPI: B04-B04A; B04-B04C; B12-A01; B12-J04; B12-K04;
     EPI: S03-E14H4
=> d all 161 tot
L61 ANSWER 1 OF 5 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
     2000-038242 [03] WPIX
AN
     1993-093726 [11]; 2000-012250 [01]
    C2000-009691
DNC
     Purified Moraxella catarrhalis outer membrane proteins useful
     for vaccinating against chronic otis media, acute maxillary sinusitis and
     other bronchopulmonary and lower respiratory tract infections.
DC
     B04 D16
     HANSEN, E J; HELMINEN, M E; MACIVER, I (TEXA) UNIV TEXAS
IN
PA
CYC
                     A 19991130 (200003)*
PΙ
     US 5993826
                                                  50
                                                        A61K039-102
ADT
     US 5993826 A CIP of US 1991-745591 19910815, CIP of WO 1992-US6869
     19920814, US 1993-25363 19930302
     US 5993826 A CIP of US 5552146
FDT
PRAI US 1993-25363
                           19930302; US 1991-745591
                                                           19910815:
     WO 1992-US6869
                           19920814
IC
     ICM A61K039-102
     ICS A61K039-02; C07K014-285; C07K016-102
AΒ
          5993826 A UPAB: 20000925
     NOVELTY - A purified Moraxella catarrhalis (also called
     Branhamella catarrhalis and Neisseria catarrhalis) 80 kiloDalton (kD) CopB
     outer membrane protein (I), is new.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:
          (i) an antigen composition (II) prepared by:
     (1) introducing a recombinant expression vector including a DNA
segment encoding (I) into a recombinant host cell;
          (2) culturing the host cell under suitable conditions for the
     expression of (I); and
          (3) collecting the expressed antigen; and
          (ii) a method (III) for inducing an antibody response to M.
     catarrhalis 80 kD CopB antigens in an animal, comprising administering
          ACTIVITY - Auditory; Respiratory active.
     MECHANISM OF ACTION - Vaccine, administration of (I) stimulates an immune response against M. catarrhalis antigens in a patient.
```

Groups of mice were immunized with the 8B6 monoclonal antibody, specific for the 80 kD outer membrane protein of M. catarrhalis. Control mice were immunized with an irrelevant antibody, 2H11 which is specific for Haemophilus ducreyi. Doses of 150 micrograms were used 18 hours prior to bacterial challenge. 5 Microliter doses of bacterial suspension, containing M. catarrhalis strain 035E, were inoculated into the lungs of the mice. 6 Hours after inoculation, the mice were sacrificed and the number of bacteria remaining in the lungs was determined. It was found that where the 2H11 antibody was used, 97% of the initial bacterial population remained. However, just 38% remained when the 8B6 antibody was used. USE - (I) may be used to vaccinate against M. catarrhalis, a pathogen implicating in causing chronic otis media, acute maxillary sinusitis and other bronchopulmonary and lower respiratory tract infections. Dwg.0/13 CPI FS FA AB; DCN MC CPI: B04-B04C1; B04-C01G; B04-E03F; B04-F0100E; B04-F10A5; B04-G09; B04-N03A; B11-C07A; B11-C08E1; B11-C09; B12-M07; B12-M08; B14-A01A5; B14-K01; B14-N02; B14-N04; B14-N05; B14-S11B; D05-C12; D05-H04; D05-H07; D05-H08; D05-H11; D05-H12A; D05-H14; D05-H17A5; D05-H18 L61 ANSWER 2 OF 5 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN 1999-560502 [47] WPIX 1990-304862 [40]; 1997-131755 [12]; 1998-413151 [35]; 2002-664558 [71] AN CR DNC C1999-163303 Isolated Haemophilus influenzae protein e useful for producing vaccines against meningitis, pneumonia, bacteremia, postpartum sepsis, acute febril tracheo-bronchitis and otitis media. DC IN GREEN, B A; ZLOTNICK, G W (PRAX-N) PRAXIS BIOLOGICS INC PA CYC PΙ US 5955580 A 19990921 (199947)\* 23 C07K001-00 US 5955580 A CIP of US 1989-320971 19890309, Div ex US 1990-491466 ADT 19900309, US 1995-449406 19950523 FDT US 5955580 A Div ex US 5601831 PRAI US 1990-491466 19900309; US 1989-320971 19890309: US 1995-449406 19950523 ICM C07K001-00 ICS A61K039-102; C07K014-285 AB US 5955580 A UPAB: 20021108 NOVELTY - Isolated Haemophilus influenzae protein e, purified of endotoxins, is new. DETAILED DESCRIPTION - An isolated protein e from Haemophilus influenzae (free from endotoxic contamination). The protein (when administered to a mammal) is capable of raising antibodies in the mammal which are protective in the infant rat passive immunization model. USE - The proteins can be used for vaccination against nontypable and typable H. influenzae. They can be used to immunize against diseases including meningitis, pneumonia, bacteremia, postpartum sepsis, acute febrile tracheo-bronchitis or otitis media. The bactericidal antibodies induced by protein e epitopes can be used to passively immunize an individual against H. influenzae. The antibody products can also be used for the detection of e proteins (e.g. via enzyme linked immunoabsorbant assay (ELISA)) and for the diagnosis of H. influenzae disease. Dwg.0/8 FS CPI FA AB: DCN MC CPI: B04-B04C1; B04-B04M; B04-C01; B04-E03F; B04-F0100E; B04-F10A; B04-G07; B04-N0300E; B11-A; B11-C07A; B11-C08D; B11-C08E1; B11-C09; B12-K04A4; B12-K04E; B12-M05; B14-A01A; B14-C03; B14-G01; B14-G03; B14-K01; B14-L06; B14-N02; B14-N16; B14-P01; B14-S11B; D05-A01A4; D05-A01B; D05-C12; D05-H04; D05-H07; D05-H09; D05-H11; D05-H12A; D05-H17A5; D05-H18 L61 ANSWER 3 OF 5 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN WPTX AN 1999-517930 [43] CR 1996-010692 [01] C1999-151167 Antigenic peptide, oligopeptide and protein useful as vaccine for Moraxella catarrhalis. DC B04 D16 IN MURPHY, T F (UYNY) UNIV NEW YORK STATE RES FOUND

```
CYC 1
                     A 19990907 (199943)*
                                                  20
                                                         A61K039-02
PΙ
     US 5948412
    US 5948412 A CIP of US 1994-245758 19940517, US 1997-810655 19970303
    US 5948412 A CIP of US 5607846
FDT
                           19970303; US 1994-245758
PRAI US 1997-810655
     ICM A61K039-02
     ICS C07K014-00
          5948412 A UPAB: 19991124
AB
     HS
     NOVELTY - Pure antigenic peptide, oligopeptide or protein (I) with one or
     more epitopes of E, an outer membrane protein of Moraxella catarrhalis is
          DETAILED DESCRIPTION - E has an apparent molecular weight of
     35000-50000 daltons by sodium dodecyl sulfate-polyacrylamide gel
     electrophoresis (SDS-PAGE) and amino acid residues 26-429 of a defined
     sequence of 459 amino acids given in the specification. E is a
     heat-modifiable protein.
          An INDEPENDENT CLAIM is also included for an antigenic formulation
     comprising a pure peptide, oligopeptide or protein with one or more
     epitopes of E.
          ACTIVITY - Antibacterial.
          MECHANISM OF ACTION - Vaccine.
          USE - (I) can be used as immunogens in prophylactic and/or
     therapeutic vaccine formulations for active immunization and for
     generating protein-specific and peptide-specific antisera useful for
     passive immunization.
          Antigenic formulations comprising (I) can be used to prevent otitis
     media, sinusitis, conjunctivitis and lower respiratory tract infections
     caused by Moraxella catarrhalis. (I) can be used as antigens for
     diagnostic immunoassays.
     Dwg.0/3
     CPI
FΑ
     AB: DCN
     CPI: B04-B04C2; B04-B04D5; B04-C01G; B04-E03F; B04-E08;
MC
          B04-F0100E; B04-N03A; B04-N04B0E; B11-C07A4; B12-K04A4;
          B14-A01A; B14-K01; B14-N02; B14-N03;
          B14-S11B; D05-C12; D05-H07; D05-H09; D05-H11; D05-H12A;
          D05-H12D5; D05-H12E; D05-H14; D05-H17A5; D05-H18; D05-H19
L61 ANSWER 4 OF 5 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
     1986-208380 [32]
AN
DNC C1986-089596
     Oral vaccine for prophylaxis of periodontitis - comprises vaccine as
TI
     antigen containing whole cell, pilus or extract of periodontitis causing
     bacterium.
DC
     B04 D16 D21
     (LIOY) LION CORP
PA
CYC
                      A 19860627 (198632)*
PΙ
     JP 61140527
                     B2 19940817 (199431)
     JP 06062431
                                                         A61K039-02
     JP 61140527 A JP 1984-263874 19841214; JP 06062431 B2 JP 1984-263874
ADT
     19841214
FDT
     JP 06062431 B2 Based on JP 61140527
PRAI JP 1984-263874
                           19841214
IC
     A61K039-02
     ICM A61K039-02
     ICS A61K039-114
     JP 61140527 A UPAB: 19930922
     Oral vaccine for prophylaxis of periodontitis, where the vaccine is made
     as antigen, which is whole cell, pilus or extract of periodontitis causative bacterium. Bacterium is eq. Bacterioides gingivalis or
     Actinomyces viscosus.
          USE/ADVANTAGE - By oral administration of this vaccine, local
     immunity mechanism is stimulated, by antibody eq. IgA, IgM, injection by
     periodontitis causative bacterium is protected specifically. Especially
     inoculation of vaccine at juvenile age, gives long period of immunological
     competence, it is effective for prophylaxis of adult periodontitis. It is
     more safety than injective administration.
          Gingivalis 381 strain is cultured on hemin and menadione added Todd.
     Hewlet broth for 2 days. Cell is collected by centrifugation (8,000 r.p.m 15 min), washed by phosphate buffer (5 mM, pH 7.4), treated by 0.5%
     formalin over night, and inactivated vaccine of cell antigen is obtd. The
     antigen is stored in refrigerator (at -80 deg.C), and used by thawing. For
     administration, in the case of antigen, cell containing solution adjusted at 10
     power 4 - 10 power 10 /ml is administered (p.o) at 0.1-10 ml/day for 3-15
     days. In the case of pilus or extract antigen, these containing solution adjusted at 0.01-10 mg/ml is administered (p.o) at 0.1-10 ml for 3-15 days
```

```
continuously.
     0/0
FS
    CPI
FA
    AB
    CPI: B02-V02; B04-B02B1; B04-B04C1; B12-A01;
MC
          B12-L03; B12-L04; D05-H07
L61 ANSWER 5 OF 5 WPIX COPYRIGHT 2004 THE THOMSON CORP on STN
    1985-236083 [38] WPIX
DNC C1985-102472
    Vaccine against infectious bovine keratoconjunctivitis - comprises
TI
    Neisseria or Branhamella gram negative cocci.
DC
    B04 C03 D16
    (GWIN-I) GWIN R M
PA
CYC
    1
    US 4539201
                    A 19850903 (198538)*
                                                10
PΙ
ADT US 4539201 A US 1983-546600 19831028
PRAI US 1983-546600
                        19831028
IC A61K039-09; C12R001-36
         4539201 A UPAB: 19930925
AB
    US
     Medicament inducing immunity to infections bovine keratoconjunctivitis
     (IBK) in cattle comprises an effective amount of gram -ve cocci from
     Neisseria and Branhamella sp., pref. those which are nonpathogenic in
     cattle, and not N gonorrhoeae or N. meningitidis. Admin. is pref.
     topically to the eye.

ADVANTAGE - The microorganisms used are effective immune stimulators,
     producing antibodies effective to produce immunity against Moraxella
     bovis, vaccines containing which do not produce practical protection against
     IBK (pinkeye).
     0/7
FS
    CPI
FA
    AB
     CPI: B02-V; B12-A01; B12-L04; B12-L09; C02-V;
MC.
          C12-A01; C12-L04; C12-L09; D05-H07
=> b home
FILE 'HOME' ENTERED AT 13:11:19 ON 16 DEC 2004
```